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(54) Title: COMPOUNDS FOR THERAPY AND DIAGNOSIS OF LUNG CANCER AND METHODS FOR THEIR USE

(57) Abstract

Compounds and methods for treating lung cancer are provided. The inventive compounds include polypeptides containing at least a portion of a lung tumor protein. Vaccines and pharmaceutical compositions for immunotherapy of lung cancer comprising such polypeptides, or polynucleotides encoding such polypeptides, are also provided, together with polynucleotides for preparing the inventive polypeptides.

COMPOUNDS FOR THERAPY AND DIAGNOSIS OF LUNG CANCER AND METHODS FOR THEIR USE

TECHNICAL FIELD

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The present invention relates generally to compositions and methods for the treatment of lung cancer. The invention is more specifically related to nucleotide sequences that are preferentially expressed in lung tumor tissue, together with polypeptides encoded by such nucleotide sequences. The inventive nucleotide sequences and polypeptides may be used in vaccines and pharmaceutical compositions for the treatment of lung cancer.

BACKGROUND OF THE INVENTION

Lung cancer is the primary cause of cancer death among both men and women in the U.S., with an estimated 172,000 new cases being reported in 1994. The five-year survival rate among all lung cancer patients, regardless of the stage of disease at diagnosis, is only 13%. This contrasts with a five-year survival rate of 46% among cases detected while the disease is still localized. However, only 16% of lung cancers are discovered before the disease has spread.

Early detection is difficult since clinical symptoms are often not seen until the disease has reached an advanced stage. Currently, diagnosis is aided by the use of chest x-rays, analysis of the type of cells contained in sputum and fiberoptic examination of the bronchial passages. Treatment regimens are determined by the type and stage of the cancer, and include surgery, radiation therapy and/or chemotherapy. In spite of considerable research into therapies for the disease, lung cancer remains difficult to treat.

Accordingly, there remains a need in the art for improved vaccines, treatment methods and diagnostic techniques for lung cancer.

SUMMARY OF THE INVENTION

Briefly stated, the present invention provides compounds and methods for the therapy of lung cancer. In a first aspect, isolated polynucleotides encoding lung tumor polypeptides are provided, such polynucleotides comprising a nucleotide sequence selected

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herein; and (b) detecting in the sample a protein or polypeptide that binds to the binding agent. In preferred embodiments, the binding agent is an antibody, most preferably a monoclonal antibody.

In related aspects, methods are provided for monitoring the progression of lung cancer in a patient, comprising: (a) contacting a biological sample obtained from a patient with a binding agent that is capable of binding to one of the polypeptides disclosed herein; (b) determining in the sample an amount of a protein or polypeptide that binds to the binding agent; (c) repeating steps (a) and (b); and comparing the amounts of polypeptide detected in steps (b) and (c).

Within related aspects, the present invention provides antibodies, preferably monoclonal antibodies, that bind to the inventive polypeptides, as well as diagnostic kits comprising such antibodies, and methods of using such antibodies to inhibit the development of lung cancer.

The present invention further provides methods for detecting lung cancer comprising: (a) obtaining a biological sample from a patient; (b) contacting the sample with a first and a second oligonucleotide primer in a polymerase chain reaction, at least one of the oligonucleotide primers being specific for a polynucleotide that encodes one of the polypeptides disclosed herein; and (c) detecting in the sample a DNA sequence that amplifies in the presence of the first and second oligonucleotide primers. In a preferred embodiment, at least one of the oligonucleotide primers comprises at least about 10 contiguous nucleotides of a polynucleotide comprising a sequence selected from the group consisting of SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181.

In a further aspect, the present invention provides a method for detecting lung cancer in a patient comprising: (a) obtaining a biological sample from the patient; (b) contacting the sample with an oligonucleotide probe specific for a polynucleotide that encodes one of the polypeptides disclosed herein; and (c) detecting in the sample a DNA sequence that hybridizes to the oligonucleotide probe. Preferably, the oligonucleotide probe comprises at least about 15 contiguous nucleotides of a polynucleotide comprising a sequence selected from the group consisting of SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181. In related a spects, diagnostic kits comprising the above oligonucleotide probes or primers are provided.

SEQ ID NO: 14 is the determined cDNA sequence for L355C1.cons SEQ ID NO: 15 is the determined cDNA sequence for L366C1.cons SEQ ID NO: 16 is the determined cDNA sequence for L163Cla SEQ ID NO: 17 is the determined cDNA sequence for LT86-1 SEQ ID NO: 18 is the determined cDNA sequence for LT86-2 SEQ ID NO: 19 is the determined cDNA sequence for LT86-3 SEQ ID NO: 20 is the determined cDNA sequence for LT86-4 SEQ ID NO: 21 is the determined cDNA sequence for LT86-5 SEQ ID NO: 22 is the determined cDNA sequence for LT86-6 10 SEQ ID NO: 23 is the determined cDNA sequence for LT86-7 SEQ ID NO: 24 is the determined cDNA sequence for LT86-8 SEQ ID NO: 25 is the determined cDNA sequence for LT86-9 SEQ ID NO: 26 is the determined cDNA sequence for LT86-10 SEQ ID NO: 27 is the determined cDNA sequence for LT86-11 SEQ ID NO: 28 is the determined cDNA sequence for LT86-12 SEQ ID NO: 29 is the determined cDNA sequence for LT86-13 SEQ ID NO: 30 is the determined cDNA sequence for LT86-14 SEQ ID NO: 31 is the determined cDNA sequence for LT86-15 SEQ ID NO: 32 is the predicted amino acid sequence for LT86-1" SEQ ID NO: 33 is the predicted amino acid sequence for LT86-2 SEQ ID NO: 34 is the predicted amino acid sequence for LT86-3 SEQ ID NO: 35 is the predicted amino acid sequence for LT86-4 SEQ ID NO: 36 is the predicted amino acid sequence for LT86-5 SEQ ID NO: 37 is the predicted amino acid sequence for LT86-6 SEQ ID NO: 38 is the predicted amino acid sequence for LT86-7 25 SEQ ID NO: 39 is the predicted amino acid sequence for LT86-8 SEQ ID NO: 40 is the predicted amino acid sequence for LT86-9 SEQ ID NO: 41 is the predicted amino acid sequence for LT86-10 SEQ ID NO: 42 is the predicted amino acid sequence for LT86-11 SEQ ID NO: 43 is the predicted amino acid sequence for LT86-12

SEQ ID NO: 74 is the predicted amino acid sequence for LT86-21 SEQ ID NO: 75 is the predicted amino acid sequence for LT86-22 SEQ ID NO: 76 is the predicted amino acid sequence for LT86-26 SEQ ID NO: 77 is the predicted amino acid sequence for LT86-27 SEQ ID NO: 78 is the determined extended cDNA sequence for L86S-12 SEQ ID NO: 79 is the determined extended cDNA sequence for L86S-36 SEQ ID NO: 80 is the determined extended cDNA sequence for L86S-46 SEQ ID NO: 81 is the predicted extended amino acid sequence for L86S-12 SEQ ID NO: 82 is the predicted extended amino acid sequence for L86S-36 SEQ ID NO: 83 is the predicted extended amino acid sequence for L86S-46 SEQ ID NO: 84 is the determined 5'cDNA sequence for L86S-6 SEQ ID NO: 85 is the determined 5'cDNA sequence for L86S-11 SEQ ID NO: 86 is the determined 5'cDNA sequence for L86S-14 SEQ ID NO: 87 is the determined 5'cDNA sequence for L86S-29 SEQ ID NO: 88 is the determined 5'cDNA sequence for L86S-34 SEQ ID NO: 89 is the determined 5'cDNA sequence for L86S-39 SEQ ID NO: 90 is the determined 5'cDNA sequence for L86S-47 SEQ ID NO: 91 is the determined 5'cDNA sequence for L86S-49 SEQ ID NO: 92 is the determined 5'cDNA sequence for L86S-51 SEQ ID NO: 93 is the predicted amino acid sequence for L86S-6 SEQ ID NO: 94 is the predicted amino acid sequence for L86S-11 SEQ ID NO: 95 is the predicted amino acid sequence for L86S-14 SEQ ID NO: 96 is the predicted amino acid sequence for L86S-29 SEQ ID NO: 97 is the predicted amino acid sequence for L86S-34 SEQ ID NO: 98 is the predicted amino acid sequence for L86S-39 SEQ ID NO: 99 is the predicted amino acid sequence for L86S-47 SEQ ID NO: 100 is the predicted amino acid sequence for L86S-49 SEQ ID NO: 101 is the predicted amino acid sequence for L86S-51 SEQ ID NO: 102 is the determined DNA sequence for SLT-T1 SEQ ID NO: 103 is the determined 5' cDNA sequence for SLT-T2

SEQ ID NO: 134 is the determined cDNA sequence for PSLT-69 SEQ ID NO: 135 is the determined cDNA sequence for PSLT-71 SEQ ID NO: 136 is the determined cDNA sequence for PSLT-73 SEQ ID NO: 137 is the determined cDNA sequence for PSLT-79 SEQ ID NO: 138 is the determined cDNA sequence for PSLT-03 SEQ ID NO: 139 is the determined cDNA sequence for PSLT-09 SEQ ID NO: 140 is the determined cDNA sequence for PSLT-011 SEQ ID NO: 141 is the determined cDNA sequence for PSLT-041 SEQ ID NO: 142 is the determined cDNA sequence for PSLT-62 SEQ ID NO: 143 is the determined cDNA sequence for PSLT-6 SEQ ID NO: 144 is the determined cDNA sequence for PSLT-37 SEQ ID NO: 145 is the determined cDNA sequence for PSLT-74 SEQ ID NO: 146 is the determined cDNA sequence for PSLT-010 SEQ ID NO: 147 is the determined cDNA sequence for PSLT-012 SEQ ID NO: 148 is the determined cDNA sequence for PSLT-037 SEQ ID NO: 149 is the determined 5' cDNA sequence for SAL-3 SEQ ID NO: 150 is the determined 5' cDNA sequence for SAL-24 SEQ ID NO: 151 is the determined 5' cDNA sequence for SAL-25 SEQ ID NO: 152 is the determined 5' cDNA sequence for SAL-33 SEQ ID NO: 153 is the determined 5° cDNA sequence for SAL-50 SEQ ID NO: 154 is the determined 5' cDNA sequence for SAL-57 SEQ ID NO: 155 is the determined 5' cDNA sequence for SAL-66 SEQ ID NO: 156 is the determined 5' cDNA sequence for SAL-82 SEQ ID NO: 157 is the determined 5' cDNA sequence for SAL-99 SEQ ID NO: 158 is the determined 5' cDNA sequence for SAL-104 SEQ ID NO: 159 is the determined 5' cDNA sequence for SAL-109 SEQ ID NO: 160 is the determined 5' cDNA sequence for SAL-5 SEQ ID NO: 161 is the determined 5' cDNA sequence for SAL-8 SEQ ID NO: 162 is the determined 5' cDNA sequence for SAL-12 SEQ ID NO: 163 is the determined 5' cDNA sequence for SAL-14

SEQ ID NO: 194 is the predicted amino acid sequence for SAL-5 SEQ ID NO: 195 is the predicted amino acid sequence for SAL-8 SEQ ID NO: 196 is the predicted amino acid sequence for SAL-12 SEQ ID NO: 197 is the predicted amino acid sequence for SAL-14 SEQ ID NO: 198 is the predicted amino acid sequence for SAL-16 SEQ ID NO: 199 is the predicted amino acid sequence for SAL-23 SEQ ID NO: 200 is the predicted amino acid sequence for SAL-26 SEQ ID NO: 201 is the predicted amino acid sequence for SAL-29 SEQ ID NO: 202 is the predicted amino acid sequence for SAL-32 SEQ ID NO: 203 is the predicted amino acid sequence for SAL-39 SEQ ID NO: 204 is the predicted amino acid sequence for SAL-42 SEQ ID NO: 205 is the predicted amino acid sequence for SAL-43 SEQ ID NO: 206 is the predicted amino acid sequence for SAL-44 SEQ ID NO: 207 is the predicted amino acid sequence for SAL-48 SEQ ID NO: 208 is the predicted amino acid sequence for SAL-68 SEQ ID NO: 209 is the predicted amino acid sequence for SAL-72 SEQ ID NO: 210 is the predicted amino acid sequence for SAL-77 SEQ ID NO: 211 is the predicted amino acid sequence for SAL-86 SEQ ID NO: 212 is the predicted amino acid sequence for SAL-88 SEQ ID NO: 213 is the predicted amino acid sequence for SAL-93 SEQ ID NO: 214 is the predicted amino acid sequence for SAL-100 SEQ ID NO: 215 is the predicted amino acid sequence for SAL-105 SEQ ID NO: 216 is a second predicted amino acid sequence for SAL-50

25 DETAILED DESCRIPTION OF THE INVENTION

As noted above, the present invention is generally directed to compositions and methods for the therapy of lung cancer. The compositions described herein include polypeptides, fusion proteins and polynucleotides. Also included within the present invention are molecules (such as an antibody or fragment thereof) that bind to the inventive polypeptides. Such molecules are referred to herein as "binding agents."

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of the proteins described herein may be identified in antibody binding assays. Such assays may generally be performed using any of a variety of means known to those of ordinary skill in the art, as described, for example, in Harlow and Lane, Antibodies: A Laboratory Manual, Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, 1988. For example, a polypeptide may be immobilized on a solid support (as described below) and contacted with patient sera to allow binding of antibodies within the sera to the immobilized polypeptide. Unbound sera may then be removed and bound antibodies detected using, for example, ¹²⁵I-labeled Protein A. Alternatively, a polypeptide may be used to generate monoclonal and polyclonal antibodies for use in detection of the polypeptide in blood or other fluids of lung cancer patients. Methods for preparing and identifying immunogenic portions of antigens of known sequence are well known in the art and include those summarized in Paul, Fundamental Immunology, 3rd ed., Raven Press, 1993, pp. 243-247.

The term "polynucleotide(s)," as used herein, means a single or double-stranded polymer of deoxyribonucleotide or ribonucleotide bases and includes DNA and corresponding RNA molecules, including HnRNA and mRNA molecules, both sense and anti-sense strands, and comprehends cDNA, genomic DNA and recombinant DNA, as well as wholly or partially synthesized polynucleotides. An HnRNA molecule contains introns and corresponds to a DNA molecule in a generally one-to-one manner. An mRNA molecule corresponds to an HnRNA and DNA molecule from which the introns have been excised. A polynucleotide may consist of an entire gene, or any portion thereof. Operable anti-sense polynucleotides may comprise a fragment of the corresponding polynucleotide, and the definition of "polynucleotide" therefore includes all such operable anti-sense fragments.

The compositions and methods of the present invention also encompass variants of the above polypeptides and polynucleotides.

A polypeptide "variant," as used herein, is a polypeptide that differs from the recited polypeptide only in conservative substitutions and/or modifications, such that the antigenic properties of the polypeptide are retained. In a preferred embodiment, variant polypeptides differ from an identified sequence by substitution, deletion or addition of five amino acids or fewer. Such variants may generally be identified by modifying one of the above polypeptide sequences, and evaluating the antigenic properties of the modified polypeptide using, for example, the representative procedures described herein. Polypeptide

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SSC, 0.5% SDS, 1.0 mM EDTA (pH 8.0); hybridizing at 50°C-65°C, 5X SSC, overnight or, in the event of cross-species homology, at 45°C with 0.5X SSC; followed by washing twice at 65°C for 20 minutes with each of 2X, 0.5X and 0.2X SSC containing 0.1% SDS. Such hybridizing DNA sequences are also within the scope of this invention, as are nucleotide sequences that, due to code degeneracy, encode an immunogenic polypeptide that is encoded by a hybridizing DNA sequence.

Two nucleotide or polypeptide sequences are said to be "identical" if the sequence of nucleotides or amino acid residues in the two sequences is the same when aligned for maximum correspondence as described below. Comparisons between two sequences are typically performed by comparing the sequences over a comparison window to identify and compare local regions of sequence similarity. A "comparison window" as used herein, refers to a segment of at least about 20 contiguous positions, usually 30 to about 75, 40 to about 50, in which a sequence may be compared to a reference sequence of the same number of contiguous positions after the two sequences are optimally aligned.

Optimal alignment of sequences for comparison may be conducted using the Megalign program in the Lasergene suite of bioinformatics software (DNASTAR, Inc., Madison, WI), using default parameters. This program embodies several alignment schemes described in the following references: Dayhoff, M.O. (1978) A model of evolutionary change in proteins - Matrices for detecting distant relationships. In Dayhoff, M.O. (ed.) Atlas of Protein Sequence and Structure, National Biomedical Resarch Foundation, Washington DC Vol. 5, Suppl. 3, pp. 345-358; Hein J. (1990) Unified Approach to Alignment and Phylogenes pp. 626-645 Methods in Enzymology vol. 183, Academic Press, Inc., San Diego, CA; Higgins, D.G. and Sharp, P.M. (1989) Fast and sensitive multiple sequence alignments on a microcomputer CABIOS 5:151-153; Myers, E.W. and Muller W. (1988) Optimal alignments in linear space CABIOS 4:11-17; Robinson, E.D. (1971) Comb. Theor 11:105; Santou, N. Nes, M. (1987) The neighbor joining method. A new method for reconstructing phylogenetic trees Mol. Biol. Evol. 4:406-425; Sneath, P.H.A. and Sokal, R.R. (1973) Numerical Taxonomy - the Principles and Practice of Numerical Taxonomy, Freeman Press, San Francisco, CA; Wilbur, W.J. and Lipman, D.J. (1983) Rapid similarity searches of nucleic acid and protein data banks Proc. Natl. Acad., Sci. USA 80:726-730.

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libraries prepared from SCID mice with mouse anti-tumor sera, as described below in Example 4. Examples of cDNA sequences that may be isolated using this technique are provided in SEQ ID NO: 149-181.

A gene encoding a polypeptide described herein (or a portion thereof) may, alternatively, be amplified from human genomic DNA, or from lung tumor cDNA, via polymerase chain reaction. For this approach, sequence-specific primers may be designed based on the nucleotide sequences provided herein and may be purchased or synthesized. An amplified portion of a specific nucleotide sequence may then be used to isolate the full length gene from a human genomic DNA library or from a lung tumor cDNA library, using well known techniques, such as those described in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, Cold Spring Harbor Laboratories, Cold Spring Harbor, NY (1989).

Once a DNA sequence encoding a polypeptide is obtained, the polypeptide may be produced recombinantly by inserting the DNA sequence into an expression vector and expressing the polypeptide in an appropriate host. Any of a variety of expression vectors known to those of ordinary skill in the art may be employed to express recombinant polypeptides of this invention. Expression may be achieved in any appropriate host cell that has been transformed or transfected with an expression vector containing a polynucleotide that encodes the recombinant polypeptide. Suitable host cells include prokaryotes, yeast and higher eukaryotic cells. Preferably, the host cells employed are *E. coli*, yeast or a mammalian cell line, such as COS or CHO cells. The DNA sequences expressed in this manner may encode naturally occurring polypeptides, portions of naturally occurring polypeptides, or other variants thereof. Supernatants from suitable host/vector systems which secrete the recombinant polypeptide may be first concentrated using a commercially available filter. The concentrate may then be applied to a suitable purification matrix, such as an affinity matrix or ion exchange resin. Finally, one or more reverse phase HPLC steps can be employed to further purify the recombinant polypeptide.

Such techniques may also be used to prepare polypeptides comprising portions or variants of the native polypeptides. Portions and other variants having fewer than about 100 amino acids, and generally fewer than about 50 amino acids, may be generated using techniques well known to those of ordinary skill in the art. For example, such polypeptides may be synthesized using any of the commercially available solid-phase techniques, such as

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extended conformation; (2) their inability to adopt a secondary structure that could interact with functional epitopes on the first and second polypeptides; and (3) the lack of hydrophobic or charged residues that might react with the polypeptide functional epitopes. Preferred peptide linker sequences contain Gly, Asn and Ser residues. Other near neutral amino acids, such as Thr and Ala may also be used in the linker sequence. Amino acid sequences which may be usefully employed as linkers include those disclosed in Maratea et al., Gene 40:39-46, 1985; Murphy et al., Proc. Natl. Acad. Sci. USA 83:8258-8262, 1986; U.S. Patent No. 4,935,233 and U.S. Patent No. 4,751,180. The linker sequence may be from 1 to about 50 amino acids in length. Peptide sequences are not required when the first and second polypeptides have non-essential N-terminal amino acid regions that can be used to separate the functional domains and prevent steric interference.

The ligated DNA sequences are operably linked to suitable transcriptional or translational regulatory elements. The regulatory elements responsible for expression of DNA are located only 5' to the DNA sequence encoding the first polypeptides. Similarly, stop codons require to end translation and transcription termination signals are only present 3' to the DNA sequence encoding the second polypeptide.

Fusion proteins are also provided that comprise a polypeptide of the present invention together with an unrelated immunogenic protein. Preferably the immunogenic protein is capable of eliciting a recall response. Examples of such proteins include tetanus, tuberculosis and hepatitis proteins (see, for example, Stoute et al. New Engl. J. Med., 336:86-91 (1997)).

Polypeptides that comprise an immunogenic portion of a lung tumor protein may generally be used for therapy of lung cancer, wherein the polypeptide stimulates the patient's own immune response to lung tumor cells. The present invention thus provides methods for using one or more of the compounds described herein (which may be polypeptides, polynucleotides or fusion proteins) for immunotherapy of lung cancer in a patient. As used herein, a "patient" refers to any warm-blooded animal, preferably a human. A patient may be afflicted with disease, or may be free of detectable disease. Accordingly, the compounds disclosed herein may be used to treat lung cancer or to inhibit the development of lung cancer. In a preferred embodiment, the compounds are administered

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ordinary skill in the art. The DNA may also be "naked," as described, for example, in published PCT application WO 90/11092, and Ulmer et al., Science 259:1745-1749, 1993, reviewed by Cohen, Science 259:1691-1692, 1993. The uptake of naked DNA may be increased by coating the DNA onto biodegradable beads, which are efficiently transported into the cells.

Routes and frequency of administration, as well as dosage, will vary from individual to individual and may parallel those currently being used in immunotherapy of other diseases. In general, the pharmaceutical compositions and vaccines may be administered by injection (e.g., intracutaneous, intramuscular, intravenous or subcutaneous), intranasally (e.g., by aspiration) or orally. Between 1 and 10 doses may be administered over a 3-24 week period. Preferably, 4 doses are administered, at an interval of 3 months, and booster administrations may be given periodically thereafter. Alternate protocols may be appropriate for individual patients. A suitable dose is an amount of polypeptide or DNA that is effective to raise an immune response (cellular and/or humoral) against lung tumor cells in a treated patient. A suitable immune response is at least 10-50% above the basal (i.e., untreated) level. In general, the amount of polypeptide present in a dose (or produced in situ by the DNA in a dose) ranges from about 1 pg to about 100 mg per kg of host, typically from about 10 pg to about 1 mg, and preferably from about 100 pg to about 1 µg. Suitable dose sizes will vary with the size of the patient, but will typically range from about 0.01 mL to about 5 mL.

While any suitable carrier known to those of ordinary skill in the art may be employed in the pharmaceutical compositions of this invention, the type of carrier will vary depending on the mode of administration. For parenteral administration, such as subcutaneous injection, the carrier preferably comprises water, saline, alcohol, a lipid, a wax and/or a buffer. For oral administration, any of the above carriers or a solid carrier, such as mannitol, lactose, starch, magnesium stearate, sodium saccharine, talcum, cellulose, glucose, sucrose, and/or magnesium carbonate, may be employed. Biodegradable microspheres (e.g., polylactic glycolide) may also be employed as carriers for the pharmaceutical compositions of this invention. Suitable biodegradable microspheres are disclosed, for example, in U.S. Patent Nos. 4.897,268 and 5.075,109.

(Natural Killer cells, lymphokine-activated killer cells), B cells, or antigen presenting cells (such as dendritic cells and macrophages) expressing the disclosed antigens. The polypeptides disclosed herein may also be used to generate antibodies or anti-idiotypic antibodies (as in U.S. Patent No. 4,918,164), for passive immunotherapy.

The predominant method of procuring adequate numbers of T-cells for adoptive immunotherapy is to grow immune T-cells in vitro. Culture conditions for expanding single antigen-specific T-cells to several billion in number with retention of antigen recognition in vivo are well known in the art. These in vitro culture conditions typically utilize intermittent stimulation with antigen, often in the presence of cytokines, such as IL-2, and non-dividing feeder cells. As noted above, the immunoreactive polypeptides described herein may be used to rapidly expand antigen-specific T cell cultures in order to generate sufficient number of cells for immunotherapy. In particular, antigen-presenting cells, such as dendritic, macrophage or B-cells, may be pulsed with immunoreactive polypeptides or transfected with a polynucleotide sequence(s), using standard techniques well known in the art. For cultured T-cells to be effective in therapy, the cultured T-cells must be able to grow and distribute widely and to survive long term in vivo. Studies have demonstrated that cultured T-cells can be induced to grow in vivo and to survive long term in substantial numbers by repeated stimulation with antigen supplemented with IL-2 (see, for example, Cheever et al. Ibid).

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The polypeptides disclosed herein may also be employed to generate and/or isolate tumor-reactive T-cells, which can then be administered to the patient. In one technique, antigen-specific T-cell lines may be generated by *in vivo* immunization with short peptides corresponding to immunogenic portions of the disclosed polypeptides. The resulting antigen specific CD8+ CTL clones may be isolated from the patient, expanded using standard tissue culture techniques, and returned to the patient.

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Alternatively, peptides corresponding to immunogenic portions of the polypeptides may be employed to generate tumor reactive T cell subsets by selective in vitro stimulation and expansion of autologous T cells to provide antigen-specific T cells which may be subsequently transferred to the patient as described, for example, by Chang et al. (Crit. Rev. Oncol. Hematol., 22(3), 213, 1996).

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at least about 80%, and preferably at least about 90%) of the patients for which lung cancer would be indicated using the full length protein, and that indicate the absence of lung cancer in substantially all of those samples that would be negative when tested with full length protein. The representative assays described below, such as the two-antibody sandwich assay, may generally be employed for evaluating the ability of a binding agent to detect metastatic human lung tumors.

The ability of a polypeptide prepared as described herein to generate antibodies capable of detecting primary or metastatic human lung tumors may generally be evaluated by raising one or more antibodies against the polypeptide (using, for example, a representative method described herein) and determining the ability of such antibodies to detect such tumors in patients. This determination may be made by assaying biological samples from patients with and without primary or metastatic lung cancer for the presence of a polypeptide that binds to the generated antibodies. Such test assays may be performed, for example, using a representative procedure described below. Polypeptides that generate antibodies capable of detecting at least 20% of primary or metastatic lung tumors by such procedures are considered to be useful in assays for detecting primary or metastatic human lung tumors. Polypeptide specific antibodies may be used alone or in combination to improve sensitivity.

Polypeptides capable of detecting primary or metastatic human lung tumors 20 may be used as markers for diagnosing lung cancer or for monitoring disease progression in patients. In one embodiment, lung cancer in a patient may be diagnosed by evaluating a biological sample obtained from the patient for the level of one or more of the above polypeptides, relative to a predetermined cut-off value. As used herein, suitable "biological samples" include blood, sera, urine and/or lung secretions.

The level of one or more of the above polypeptides may be evaluated using any binding agent specific for the polypeptide(s). A "binding agent," in the context of this invention, is any agent (such as a compound or a cell) that binds to a polypeptide as described above. As used herein, "binding" refers to a noncovalent association between two separate molecules (each of which may be free (i.e., in solution) or present on the surface of a cell or a solid support), such that a "complex" is formed. Such a complex may be free or immobilized (either covalently or noncovalently) on a support material. The ability to bind may generally

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be immobilized on the solid support using a variety of techniques known to those of skill in the art, which are amply described in the patent and scientific literature. In the context of the present invention, the term "immobilization" refers to both noncovalent association, such as adsorption, and covalent attachment (which may be a direct linkage between the antigen and functional groups on the support or may be a linkage by way of a cross-linking agent). Immobilization by adsorption to a well in a microtiter plate or to a membrane is preferred. In such cases, adsorption may be achieved by contacting the binding agent, in a suitable buffer, with the solid support for a suitable amount of time. The contact time varies with temperature, but is typically between about 1 hour and about 1 day. In general, contacting a well of a plastic microtiter plate (such as polystyrene or polyvinylchloride) with an amount of binding agent ranging from about 10 ng to about 10 µg, and preferably about 100 ng to about 1 µg, is sufficient to immobilize an adequate amount of binding agent.

Covalent attachment of binding agent to a solid support may generally be achieved by first reacting the support with a bifunctional reagent that will react with both the support and a functional group, such as a hydroxyl or amino group, on the binding agent. For example, the binding agent may be covalently attached to supports having an appropriate polymer coating using benzoquinone or by condensation of an aldehyde group on the support with an amine and an active hydrogen on the binding partner (see, e.g., Pierce Immunotechnology Catalog and Handbook, 1991, at A12-A13).

In certain embodiments, the assay is a two-antibody sandwich assay. This assay may be performed by first contacting an antibody that has been immobilized on a solid support, commonly the well of a microtiter plate, with the sample, such that polypeptides within the sample are allowed to bind to the immobilized antibody. Unbound sample is then removed from the immobilized polypeptide-antibody complexes and a second antibody (containing a reporter group) capable of binding to a different site on the polypeptide is added. The amount of second antibody that remains bound to the solid support is then determined using a method appropriate for the specific reporter group.

More specifically, once the antibody is immobilized on the support as described above, the remaining protein binding sites on the support are typically blocked. Any suitable blocking agent known to those of ordinary skill in the art, such as bovine serum albumin or Tween 20TM (Sigma Chemical Co., St. Louis, MO). The immobilized antibody is

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that corresponds to a predetermined cut-off value. In one preferred embodiment, the cut-off value is the average mean signal obtained when the immobilized antibody is incubated with samples from patients without lung cancer. In general, a sample generating a signal that is three standard deviations above the predetermined cut-off value is considered positive for lung cancer. In an alternate preferred embodiment, the cut-off value is determined using a Receiver Operator Curve, according to the method of Sackett et al., Clinical Epidemiology: A Basic Science for Clinical Medicine, Little Brown and Co., 1985, p. 106-7. Briefly, in this embodiment, the cut-off value may be determined from a plot of pairs of true positive rates (i.e., sensitivity) and false positive rates (100%-specificity) that correspond to each possible cut-off value for the diagnostic test result. The cut-off value on the plot that is the closest to the upper left-hand corner (i.e., the value that encloses the largest area) is the most accurate cut-off value, and a sample generating a signal that is higher than the cut-off value determined by this method may be considered positive. Alternatively, the cut-off value may be shifted to the left along the plot, to minimize the false positive rate, or to the right, to minimize the false negative rate. In general, a sample generating a signal that is higher than the cut-off value determined by this method is considered positive for lung cancer.

In a related embodiment, the assay is performed in a flow-through or strip test format, wherein the antibody is immobilized on a membrane, such as nitrocellulose. In the flow-through test, polypeptides within the sample bind to the immobilized antibody as the sample passes through the membrane. A second, labeled antibody then binds to the antibody-polypeptide complex as a solution containing the second antibody flows through the membrane. The detection of bound second antibody may then be performed as described above. In the strip test format, one end of the membrane to which antibody is bound is immersed in a solution containing the sample. The sample migrates along the membrane through a region containing second antibody and to the area of immobilized antibody. Concentration of second antibody at the area of immobilized antibody indicates the presence of lung cancer. Typically, the concentration of second antibody at that site generates a pattern, such as a line, that can be read visually. The absence of such a pattern indicates a negative result. In general, the amount of antibody immobilized on the membrane is selected to generate a visually discernible pattern when the biological sample contains a level of polypeptide that would be sufficient to generate a positive signal in the two-antibody

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of immortal cell lines capable of producing antibodies having the desired specificity (i.e., reactivity with the polypeptide of interest). Such cell lines may be produced, for example, from spleen cells obtained from an animal immunized as described above. The spleen cells are then immortalized by, for example, fusion with a myeloma cell fusion partner, preferably one that is syngeneic with the immunized animal. A variety of fusion techniques may be employed. For example, the spleen cells and myeloma cells may be combined with a nonionic detergent for a few minutes and then plated at low density on a selective medium that supports the growth of hybrid cells, but not myeloma cells. A preferred selection technique uses HAT (hypoxanthine, aminopterin, thymidine) selection. After a sufficient time, usually about 1 to 2 weeks, colonies of hybrids are observed. Single colonies are selected and tested for binding activity against the polypeptide. Hybridomas having high reactivity and specificity are preferred.

Monoclonal antibodies may be isolated from the supernatants of growing hybridoma colonies. In addition, various techniques may be employed to enhance the yield, such as injection of the hybridoma cell line into the peritoneal cavity of a suitable vertebrate host, such as a mouse. Monoclonal antibodies may then be harvested from the ascites fluid or the blood. Contaminants may be removed from the antibodies by conventional techniques, such as chromatography, gel filtration, precipitation, and extraction. The polypeptides of this invention may be used in the purification process in, for example, an affinity chromatography step.

Monoclonal antibodies of the present invention may also be used as therapeutic reagents, to diminish or eliminate lung tumors. The antibodies may be used on their own (for instance, to inhibit metastases) or coupled to one or more therapeutic agents. Suitable agents in this regard include radionuclides, differentiation inducers, drugs, toxins, and derivatives thereof. Preferred radionuclides include ⁹⁰Y, ¹²³I, ¹²⁵I, ¹³¹I, ¹²⁶Re, ¹²⁸Re, ²¹¹At, and ²¹²Bi. Preferred drugs include methotrexate, and pyrimidine and purine analogs. Preferred differentiation inducers include phorbol esters and butyric acid. Preferred toxins include ricin, abrin, diptheria toxin, cholera toxin, gelonin, Pseudomonas exotoxin, Shigella toxin, and pokeweed antiviral protein.

A therapeutic agent may be coupled (e.g., covalently bonded) to a suitable monoclonal antibody either directly or indirectly (e.g., via a linker group). A direct reaction

be prepared in a variety of ways. For example, more than one agent may be coupled directly to an antibody molecule, or linkers which provide multiple sites for attachment can be used. Alternatively, a carrier can be used.

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A carrier may bear the agents in a variety of ways, including covalent bonding either directly or via a linker group. Suitable carriers include proteins such as albumins (e.g., U.S. Patent No. 4,507,234, to Kato et al.), peptides and polysaccharides such as aminodextran (e.g., U.S. Patent No. 4,699,784, to Shih et al.). A carrier may also bear an agent by noncovalent bonding or by encapsulation, such as within a liposome vesicle (e.g., U.S. Patent Nos. 4,429,008 and 4,873,088). Carriers specific for radionuclide agents include radiohalogenated small molecules and chelating compounds. For example, U.S. Patent No. 4,735,792 discloses representative radiohalogenated small molecules and their synthesis. A radionuclide chelate may be formed from chelating compounds that include those containing nitrogen and sulfur atoms as the donor atoms for binding the metal, or metal oxide, radionuclide. For example, U.S. Patent No. 4,673,562, to Davison et al. discloses representative chelating compounds and their synthesis.

A variety of routes of administration for the antibodies and immunoconjugates may be used. Typically, administration will be intravenous, intramuscular, subcutaneous or in the bed of a resected tumor. It will be evident that the precise dose of the antibody/immunoconjugate will vary depending upon the antibody used, the antigen density on the tumor, and the rate of clearance of the antibody.

Diagnostic reagents of the present invention may also comprise DNA sequences encoding one or more of the above polypeptides, or one or more portions thereof. For example, at least two oligonucleotide primers may be employed in a polymerase chain reaction (PCR) based assay to amplify lung tumor-specific cDNA derived from a biological sample, wherein at least one of the oligonucleotide primers is specific for a polynucleotide encoding a lung tumor protein of the present invention. The presence of the amplified cDNA is then detected using techniques well known in the art, such as gel electrophoresis. Similarly, oligonucleotide probes specific for a polynucleotide encoding a lung tumor protein of the present invention may be used in a hybridization assay to detect the presence of an inventive polypeptide in a biological sample.

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The following Examples are offered by way of illustration and not by way of

EXAMPLES

Example 1

PREPARATION OF LUNG TUMOR-SPECIFIC cDNA SEQUENCES USING DIFFERENTIAL DISPLAY RT-PCR

This example illustrates the preparation of cDNA molecules encoding lung tumor-specific polypeptides using a differential display screen.

Tissue samples were prepared from breast tumor and normal tissue of a patient with lung cancer that was confirmed by pathology after removal of samples from the patient. Normal RNA and tumor RNA was extracted from the samples and mRNA was isolated and converted into cDNA using a (dT)₁₂AG (SEQ ID NO: 47) anchored 3' primer. Differential display PCR was then executed using a randomly chosen primer (SEQ ID NO: 48). Amplification conditions were standard buffer containing 1.5 mM MgCl₂, 20 pmol of primer, 500 pmol dNTP and 1 unit of Taq DNA polymerase (Perkin-Elmer, Branchburg, NJ). Forty cycles of amplification were performed using 94 °C denaturation for 30 seconds, 42 °C annealing for 1 minute and 72 °C extension for 30 seconds. Bands that were repeatedly observed to be specific to the RNA fingerprint pattern of the tumor were cut out of a silver stained gel, subcloned into the pGEM-T vector (Promega, Madison, WI) and sequenced. The isolated 3' sequences are provided in SEQ ID NO: 1-16.

Comparison of these sequences to those in the public databases using the BLASTN program, revealed no significant homologies to the sequences provided in SEQ ID NO: 1-11. To the best of the inventors' knowledge, none of the isolated DNA sequences have previously been shown to be expressed at a greater level in human lung tumor tissue than in normal lung tissue.

aminopeptidase. Clone LT86-9 appears to contain two inserts, with the 5' sequence showing homology to the previously identified antisense sequence of interferon alpha-induced P27, and the 3' sequence being similar to LT86-6. Clone LT86-14 (SEQ ID NO: 30) was found to show some homology to the trithorax gene and has an "RGD" cell attachment sequence and a beta-Lactamase A site which functions in hydrolysis of penicillin. Clones LT86-1, LT86-2. LT86-4, LT86-5 and LT86-10 (SEQ ID NOS: 17, 18, 20, 21 and 26, respectively) were found to show homology to previously identified genes. A subsequently determined extended cDNA sequence for LT86-4 is provided in SEQ ID NO: 66, with the corresponding predicted amino acid sequence being provided in SEQ ID NO: 67.

Subsequent studies led to the isolation of five additional clones, referred to as LT86-20, LT86-21, LT86-22, LT86-26 and LT86-27. The determined 5' cDNA sequences for LT86-20, LT86-22, LT86-26 and LT86-27 are provided in SEQ ID NO: 68 and 70-72, respectively, with the determined 3' cDNA sequences for LT86-21 being provided in SEQ ID NO: 69. The corresponding predicted amino acid sequences for LT86-20, LT86-21, LT86-22, LT86-26 and LT86-27 are provided in SEQ ID NO: 73-77, respectively. LT86-22 and LT86-27 were found to be highly similar to each other. Comparison of these sequences to those in the gene bank as described above, revealed no significant homologies to LT86-22 and LT86-27. LT86-20, LT86-21 and LT86-26 were found to show homology to previously identified genes.

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predicted amino acid sequences are provided in SEQ ID NO: 93-101, respectively. L86S-30, L86S-39 and L86S-47 were found to be similar to each other. Comparison of these sequences with those in the gene bank as described above, revealed no significant homologies to L86S-14. L86S-29 was found to show some homology to a previously identified EST. L86S-6, L86S-11, L86S-34, L86S-39, L86S-47, L86S-49 and L86S-51 were found to show some homology to previously identified genes.

In further studies, a directional cDNA library was constructed using a Stratagene kit with a Lambda Zap Express vector. Total RNA for the library was isolated from two primary squamous lung tumors and poly A+RNA was isolated using an oligo dT column. Antiserum was developed in normal mice using a pool of sera from three SCID mice implanted with human squamous lung carcinomas. Approximately 700,000 PFUs were screened from the unamplified library with E. coli absorbed mouse anti-SCID tumor serum. Positive plaques were identified as described above. Phage was purified and phagemid excised for 180 clones with inserts in a pBK-CMV vector for expression in prokaryotic or eukaryotic cells.

The determined cDNA sequences for 23 of the isolated clones are provided in SEQ ID NO: 126-148. Comparison of these sequences with those in the public database as described above revealed no significant homologies to the sequences of SEQ ID NO: 139 and 143-148. The sequences of SEQ ID NO: 126-138 and 140-142 were found to show homology previously identified human polynucleotide sequences.

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tags (ESTs). The sequences of SEQ ID NO: 150, 155 and 159-181 were found to show homology to sequences previously identified in humans.

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Example 6

ISOLATION OF DNA SEQUENCES ENCODING LUNG TUMOR ANTIGENS

DNA sequences encoding antigens potentially involved in squamous cell lung tumor formation were isolated as follows.

A lung tumor directional cDNA expression library was constructed employing the Lambda ZAP Express expression system (Stratagene, La Jolla, CA). Total RNA for the library was taken from a pool of two human squamous epithelial lung carcinomas and poly A+ RNA was isolated using oligo-dT cellulose (Gibco BRL, Gaithersburg, MD). Phagemid were rescued at random and the cDNA sequences of isolated clones were determined.

The determined cDNA sequence for the clone SLT-T1 is provided in SEQ ID NO: 102, with the determined 5' cDNA sequences for the clones SLT-T2, SLT-T3, SLT-T5, SLT-T7, SLT-T9, SLT-T10, SLT-T11 and SLT-T12 being provided in SEQ ID NO: 103-110, respectively. The corresponding predicted amino acid sequence for SLT-T1, SLT-T2, SLT-T3, SLT-T10 and SLT-T12 are provided in SEQ ID NO: 111-115, respectively. Comparison of the sequences for SLT-T2, SLT-T3, SLT-T5, SLT-T7, SLT-T9 and SLT-T11 with those in the public databases as described above, revealed no significant homologies. The sequences for SLT-T10 and SLT-T12 were found to show some homology to sequences previously identified in humans.

The sequence of SLT-T1 was determined to show some homology to a PAC clone of unknown protein function. The cDNA sequence of SLT-T1 (SEQ ID NO: 102) was found to contain a mutator (MUTT) domain. Such domains are known to function in removal of damaged guanine from DNA that can cause A to G transversions (see, for example, el-Deiry, W.S., 1997 Curr. Opin. Oncol. 9:79-87; Okamoto, K. et al. 1996 Int. J. Cancer 65:437-41; Wu, C. et al. 1995 Biochem. Biophys. Res. Commun. 214:1239-45; Porter, D.W. et al. 1996 Chem. Res. Toxicol. 9:1375-81). SLT-T1 may thus be of use in the treatment, by gene therapy, of lung cancers caused by, or associated with, a disruption in DNA repair.

Example 7 SYNTHESIS OF POLYPEPTIDES

Polypeptides may be synthesized on a Perkin Elmer/Applied Biosystems

Division 430A peptide synthesizer using FMOC chemistry with HPTU (O-BenzotriazoleN,N,N',N'-tetramethyluronium hexafluorophosphate) activation. A Gly-Cys-Gly sequence
may be attached to the amino terminus of the peptide to provide a method of conjugation,
binding to an immobilized surface, or labeling of the peptide. Cleavage of the peptides from
the solid support may be carried out using the following cleavage mixture: trifluoroacetic

acid:ethanedithiol:thioanisole:water:phenol (40:1:2:2:3). After cleaving for 2 hours, the
peptides may be precipitated in cold methyl-t-butyl-ether. The peptide pellets may then be
dissolved in water containing 0:1% trifluoroacetic acid (TFA) and lyophilized prior to
purification by C18 reverse phase HPLC. A gradient of 0%-60% acetonitrile (containing
0.1% TFA) in water (containing 0.1% TFA) may be used to elute the peptides. Following
lyophilization of the pure fractions, the peptides may be characterized using electrospray or
other types of mass spectrometry and by amino acid analysis.

From the foregoing, it will be appreciated that, although specific embodiments of the invention have been described herein for the purposes of illustration, various modifications may be made without deviating from the spirit and scope of the invention.

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- A vaccine comprising the polypeptide of claim 2 and an immune response enhancer.
- 5 10. The vaccine of claim 9 wherein the immune response enhancer is an adjuvant.
 - 11. A vaccine comprising the polynucleotide of claims 1 or 4 and an immune response enhancer.
 - 12. The vaccine of claim 11 wherein the immune response enhancer is an adjuvant.
- 13. A pharmaceutical composition for the treatment of lung cancer comprising a polypeptide and a physiologically acceptable carrier, the polypeptide comprising an immunogenic portion of a lung protein or of a variant thereof, wherein said protein comprises an amino acid sequence encoded by a polynucleotide comprising a sequence selected from the group consisting of:
 - (a) sequences recited in SEQ ID NO: 12-18, 20, 21, 26, 49, 50, 52, 54, 64, 66, 68, 69, 71, 78, 84, 85, 88, 91, 92, 116-120, 126-138, 140-142, 150, 155 and 159-181;
 - (b) sequences complementary to the sequences of SEQ ID NO: 12-18, 20, 21, 26, 49, 50, 52, 54, 64, 66, 68, 69, 71, 78, 84, 85, 88, 91, 92, 116-120, 126-138, 140-142, 150, 155 and 159-181; and
 - (c) variants of the sequences of (a) and (b).

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- 14. A vaccine for the treatment of lung cancer comprising a polypeptide and an immune response enhancer, said polypeptide comprising an immunogenic portion of a lung protein or of a variant thereof, wherein said protein comprises an amino acid sequence encoded by a polynucleotide comprising a sequence selected from the group consisting of:
- (a) sequences recited in SEQ ID NO: 12-18, 20, 21, 26, 49, 50, 52, 54, 64, 66, 68, 69, 71, 78, 84, 85, 88, 91, 92, 116-120, 126-138, 140-142, 150, 155 and 159-181;

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- 21. A pharmaceutical composition comprising a fusion protein according to any one of claims 18-20 and a physiologically acceptable carrier.
- 5 22. A vaccine comprising a fusion protein according to any one of claims 18-20 and an immune response enhancer.
 - 23. The vaccine of claim 22 wherein the immune response enhancer is an adjuvant.
 - 24. A method for inhibiting the development of lung cancer in a patient, comprising administering to the patient an effective amount of the pharmaceutical composition of claim 21.
 - 25. A method for inhibiting the development of lung cancer in a patient, comprising administering to the patient an effective amount of the vaccine of claim 22.
 - 26. A method for inhibiting the development of lung cancer in a patient, comprising administering to the patient a polynucleotide under conditions such that the polynucleotide enters a cell of the patient and is expressed therein, the polynucleotide having a sequence selected from the group consisting of:
 - (a) a sequence provided in SEQ ID NO: 102;
 - (b) sequences complementary to a sequence of SEQ ID NO: 102; and
 - (c) variants of the sequence of SEQ ID NO: 102.
 - 27. A method for detecting lung cancer in a patient, comprising:
 - (a) contacting a biological sample obtained from the patient with a binding agent which is capable of binding to a polypeptide, the polypeptide comprising an immunogenic portion of a lung tumor protein or a variant thereof, wherein said protein comprises an amino acid sequence encoded by a polynucleotide comprising a nucleotide sequence selected from the group consisting of sequences provided in SEQ ID NO: 1-31, 49-

- (a) sequences recited in SEQ ID NO: 1-11, 19, 22-25, 27-31, 51, 53, 55, 63, 70, 72, 79, 80, 86, 87, 89, 90, 102-107, 109, 139, 143-149, 151-154 and 156-158;
- (b) the complements of nucleotide sequences recited in SEQ ID NO: 1-11, 19, 22-25, 27-31, 51, 53, 55, 63, 70, 72, 79, 80, 86, 87, 89, 90, 102-107, 109, 139, 143-149, 151-154 and 156-158; and
- (c) variants of the sequences of (a) and (b).
- 32. A method for inhibiting the development of lung cancer in a patient, comprising administering to the patient a therapeutically effective amount of a monoclonal antibody according to claim 31.
 - 33. The method of claim 32 wherein the monoclonal antibody is conjugated to a therapeutic agent.
 - 34. A method for detecting lung cancer in a patient comprising:
 - (a) obtaining a biological sample from the patient;
- (b) contacting the sample with at least two oligonucleotide primers in a polymerase chain reaction, wherein at least one of the oligonucleotides is specific for a polynucleotide encoding a polypeptide comprising an immunogenic portion of a lung tumor protein or a variant thereof, said protein comprising an amino acid sequence encoded by a polynucleotide comprising a nucleotide sequence selected from the group consisting of sequences recited in SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181, the complements of said sequences and variants thereof; and
 - (c) detecting in the sample a DNA sequence that amplifies in the presence of the oligonucleotide primers, thereby detecting lung cancer.
- 35. The method of claim 34, wherein at least one of the oligonucleotide primers comprises at least about 10 contiguous nucleotides of a polynucleotide comprising a sequence selected from SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181.

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provided in SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181, the complements of said sequences and variants thereof.

- 44. A method for detecting lung cancer in a patient, comprising:
- (a) obtaining a biological sample from the patient;
- (b) contacting the biological sample with an oligonucleotide probe specific for a polynucleotide encoding a polypeptide comprising an immunogenic portion of a lung tumor protein or a variant thereof, said protein comprising an amino acid sequence encoded by a polynucleotide comprising a nucleotide sequence selected from the group consisting of sequences recited in SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181, the complements of said nucleotide sequences and variants thereof; and
- (c) detecting in the sample a DNA sequence that hybridizes to the oligonucleotide probe, thereby detecting lung cancer in the patient.
- 45. The method of claim 44 wherein the oligonucleotide probe comprises at least about 15 contiguous nucleotides of a polynucleotide having a nucleotide sequence selected from the group consisting of sequences recited in SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181, the complements of said nucleotide sequences and variants thereof.
- 46. A diagnostic kit comprising an oligonucleotide probe specific for a polynucleotide encoding a polypeptide comprising an immunogenic portion of a lung tumor protein or a variant thereof, said protein comprising an amino acid sequence encoded by a polynucleotide comprising a nucleotide sequence selected from the group consisting of sequences recited in SEQ ID NO: 1-31, 49-55, 63, 64, 66, 68-72, 78-80, 84-92, 102-110, 116-120 and 126-181, the complements of said sequences and variants thereof.
- 47. The diagnostic kit of claim 46, wherein the oligonucleotide probe comprises at least about 15 contiguous nucleotides of a polynucleotide having a nucleotide sequence selected from the group consisting of sequences recited in SEQ ID NO: 1-31, 49-55,

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pharmaceutically acceptable carrier.

- 55. A composition for the treatment of lung cancer in a patient, comprising T cells proliferated in the presence of a polynucleotide of claim 1, in combination with a pharmaceutically acceptable carrier.
 - 56. A method for treating lung cancer in a patient, comprising the steps of:
 - (a) incubating antigen presenting cells in the presence of at least one polypeptide of claim 2; and
 - (b) administering to the patient the incubated antigen presenting cells.
 - 57. A method for treating lung cancer in a patient, comprising the steps of:
 - (a) incubating antigen presenting cells in the presence of at least one polynucleotide of claim 1; and
 - (b) administering to the patient the incubated antigen presenting cells.
 - 58. The method of claims 54 or 55 wherein the antigen presenting cells are selected from the group consisting of dendritic cells and macrophage cells.
 - 59. A composition for the treatment of lung cancer in a patient, comprising antigen presenting cells incubated in the presence of a polypeptide of claim 2, in combination with a pharmaceutically acceptable carrier.
- 60. A composition for the treatment of lung cancer in a patient, comprising antigen presenting cells incubated in the presence of a polynucleotide of claim 1, in combination with a pharmaceutically acceptable carrier.

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Arg Arg Glu Cys Pro Ser Asp Glu Cys Gly Ala Gly Val Phe Met Ala
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	1 .		o -	٠	5		:										
	:				,			. e 194		٠.			· 199			٠٠, ٠٠	
I	eu A	sp A	lla	Gln	Glu	His	Val	Lys	Asn	. Pro	Tyr	Lys	Gly	Lys	Lys	Leu	•
74	ta kar		₹.	× 20			- 5	74.	: 25				1.	30		. 2000	
	, <u>-</u> •			٠.			<i>3</i> .	. 4.	A 35							r gree Lyr ker	
T	we Is	ve H	lie	Pro	Asn	Phe	Pro	Lvs	INS	Pro	Leu	Thr	Pro	Tvr	Phe	Arg	٠,٠
_	·2,	,	25			4.		40					45	-3-			
٠.			.33			& 2€		- 10									
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٠.		50/.2				· .	55	A	$x = t_{T}$			60		,		ja ku	
1	2 °			•	100		2 13		3						•		
.S	er A	sn I	æu.	Asp	Leu									Lys	Glu	Leu	
٠.	65					70		*			75					80	
	1				٠	١.		(500 A)						:			
P	ro G	lu I	vs	Lvs	Lvs	Met				Pro	Asp	Phe	Gln	Arq	Arq	Glu	ŧ,
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					21-	T	Dro	C1	Dr.	T10	Cln.					Pro	
	III G.	LY A	aı.			ьys									PIO	PIO	
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₽	TO T	yr P	LO	Glu	Cys									Pro	Gln	Asp	
		1	15					120	д э , г				125	£	÷ .		
P	ro Pi	ro														**	
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S	er I	le C	P.V.	Met	Ara	Met	Glu	Ile	Leu	Glv	Cvs	Pro	Leu	Pro	Asp	Pro	
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Gly	r Glu	His 115	s Glu	va]	Gly	Glü	1 Pro		Phe	His	Tyr	11e		Gly	/ Ala
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Glu	Gly	Met	Leu 20	Met	Gly	Val	Lys	Pro 25	Gly	Glu [.]	Asp	Ala	Ser 30	Gly	Pro
Ala	Glu	Asp 35	Leu	Val	Arg	Arg	Ser 40	Glu	Lys	Asp	Thr	Ala 45	Ala	Val	Val
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Glu 65	Pro	Glu	Ala	Glu	Pro 70	Glu	Ser	Lys	Ser	Glu 75	Pro	Arg	Pro	Pro	Ile 80

Ser Ser Pro Arg Ala Pro Gln Thr Arg Ala Val Lys Pro Arg Leu His 85 90 95

Pro Val Lys Pro Met Asn Ala Thr Ala Thr Lys Val Ala Asn Cys Ser 100 105 110

Leu Gly Thr Ala Thr Ile Ile Gly Glu Asn Leu Asn Asn Glu Val Met 115 120 125

Met Lys Lys Tyr Ser Pro Ser Asp Pro Ala Phe Ala Tyr Ala Gln Leu 130 135 140

Thr His Asp Glu Leu Ile Gln Leu Val Leu Lys Gln Lys Glu Thr Ile 145 150 155 160

Ser Lys Lys Glu Phe Gln Val Arg Glu Leu Glu Asp Tyr Ile Asp Asn 165 -170 175

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Thr Gln Val Gly Lys Lys Ala Gly Lys Met 195 200

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<213> Homo sapiens

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Gly Ser Val Leu Asp Tyr Phe Ser Glu Arg Ser Asn Pro Phe Tyr Asp 35 40 45

Arg Thr Cys Asn Asn Glu Val Val Lys Met Gln Arg Leu Thr Leu Glu 50 55 60

His Leu Asn Gln Met Val Gly Ile Glu Tyr Ile Leu Leu His Ala Gln 65 70 75 80

Glu Pro Ile Leu Phe Ile Ile Arg Lys Gln Gln Arg Gln Ser Pro Ala 85 90 95

Gln Val Ile Pro Leu Ala Asp Tyr Tyr Ile Ile Ala Gly Val Ile Tyr 100 105 110

Gln Ala Pro Asp Leu Gly Ser Val Ile Asn Ser Arg Val Leu Thr Ala 115 120 125

37-3															
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Gln	Asp	Lγş	Val	Arg 165		Lys	Ala	Lys	Arg 170		Ģlu	Glu	Pro	Ser 175	Ser
Ile	Phe	Gln	Arg 180		Arg	Val	Asp	Ala 185		Leu	Leu	Asp	Leu 190	-	Gln
Lys	Ile	Ser 195		Gln	Ile	Cys	Ala 200		Asp	Gln	Thr	Lys 205		Glu	Ala
Glu	Pro 210		Pro	Glu	Thr	Val 215	Lys	Pro	Glu	Glu	Lys 220	Glu	Thr	Thr	Lys
Asn 225	Val	Gln	Gln	Thr	Val 230	Ser	Ala	Lys	Gly	Pro 235	Pro	Glu	Lys	Arg	Met 240
Arg	Leu	Gln					*			٠.				:	
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	2> P	RT	sapi	ens				•							t git Libe
<21:	2> P 3> H	RT Omo s	sapi	ens				•					-		1 (2) 1 (3)
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Phe Asp Glu Ala Met Ser Tyr Cys Arg Tyr His Pro Ser Lys Gly Tyr 130 140

Trp Trp His Phe Lys Asp His Glu Glu Gln Asp Lys Val Arg Pro Lys 145 150 155 160

Ala Lys Arg Lys Glu Glu Pro Ser Ser Ile Phe Gln Arg Gln Arg Val

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Gln Leu Lys Pro Gly Glu Lys Pro Val Pro Val Asp Gln Thr Lys Lys
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Arg Met Arg Leu Gln 245

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Pro Gly Arg Ala Pro Ala His Ala Pro Leu Ser Ser Phe Val Pro Gly 65 70 75 80

Val Gly Gly Arg Ser Pro Ala Ser Val Gly Ile Ser Ala Pro Gly Gly
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Gly Pro Ser Gly Ala Ala Ala Lys Ile Pro Leu Glu Leu Thr Gln Ser

Arg Val Gln Lys Ile Trp Val Pro Val Asp His Arg Pro Ser Leu Pro 115 120 125

Arg Ser Cys Gly Pro Lys Leu Thr Asn Ser Pro Ala Val Phe Val Met

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130		1	35				14	10			
Val Gly Le	eu Pro Ar	g Pro G 150	ly G	ln As	p Le	u Le 15		u H	is G	lu. Se	er Le 16
Leu Ala Al	la									٠.	
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<213> Homo	sapiens								,.		
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Trp Ile Pro	o Ile Let 20	ı Asn Se	er Gly	y. Ser 25	Val	Leu	Ası	ту	r Ph		r Glu
Arg Ser Ası	n Pro Phe 5	Tyr As	p Arg	Thr	Cys	Asn	Ası	1 Gl:		l Va	l Lys
Met Gln Arg	g Leu Thr	Leu Gl	u His S	Leu	Asn	Gln	Met 60		l Gly	/ Ile	e Glu
Tyr Ile Leu 65	ı Leu His	Ala Gl	n Glu	Pro	Ile	Leu 75	Phe	Ile	e Ile	. Arg	Lys 80
Gln Gln Arg	Gln Ser 85	Pro Ala	a Gln	Val	Ile 90	Pro	Leu	Ala	Asp	Ty:	
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Glu Ala Met 130	Ser Tyr	Cys Arg	Tyr	His	Pro	Ser	Lys 140	Gly	Туг	Trp	Trp
His Phe Lys 145	Asp His	Glu Glu 150	Gln	Asp	Lys	Val 155	Arg	Pro	Lys	Ala	Lys 160
Arg Lys Glu	Glu Pro 165	Ser Ser	Ile	Phe	Gln 170	Arg (G1n	Arg	Val	Asp 175	Ala
Leu Leu Leu	Asp Leu 180	Arg Gln	Lys	Phe 185	Pro 1	Pro 1	Ŀys	Phe	Val 190	Gln	Leu '
Lys Pro Gly 195	Glu Lys	Pro Val	Pro 200	Val i	Asp (3ln 1		Lys 205	Lys	Glu	Ala

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Arg Leu Gln

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Glu Arg Ser Asn Pro Phe Tyr Asp Arg Thr Cys Asn Asn Glu Val Val
35 40 45

Lys Met Gln Arg Leu Thr Leu Glu His Leu Asn Gln Met Val Gly Ile 50 60

Glu Tyr Ile Leu Leu His Ala Gln Glu Pro Ile Leu Phe Ile Ile Arg
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Lys Gln Gln Arg Gln Ser Pro Ala Gln Val Ile Pro Leu Ala Asp Tyr 85 90 95

Tyr Ile Ile Ala Gly Val Ile Tyr Gln Ala Pro Asp Leu Gly Ser Val 100 105 110

Ile Asn Ser Arg Val Leu Thr Ala Val His Gly Ile Gln Ser Ala Phe 115 120 125

Asp Glu Ala Met Ser Tyr Cys Arg Tyr His Pro Ser Lys Gly Tyr Trp 130 135 140

Trp His Phe Lys Asp His Glu Glu Gln Asp Lys Val Arg Pro Lys Ala 145 150 150 160

Lys Arg Lys Glu Glu Pro Ser Ser Ile Phe Gln Arg Gln Arg Val Asp
165 170 175

Ala Leu Leu Asp Leu Arg Gln Lys Phe Pro Pro Lys Phe Val Gln
180 185 190

Leu Lys Pro Gly Glu Lys Pro Val Pro Val Asp Gln Thr Lys Lys Glu
195 200 205

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Met Arg Leu Gln

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Val Ala Leu Leu Leu Ala Leu Phe Gly Arg Ala Ser Glu Asp Pro Leu 65 70 75 80

Leu Asn Leu Val Ser Pro Leu Asp Cys Glu Val Asp Ala Gln Glu Gly 85 90 95

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Ala Leu Cys Pro Glu Gly His Glu Trp Ser Gln Ile Tyr Phe Ser Pro 35 40 45

Ser Gly Asn Ile Val Ala His Glu Asn Cys Leu Leu Tyr Ser Ser Gly
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Leu Val Glu Cys Glu Thr Leu Asp Leu Arg Asn Thr Ile Arg Asn Phe
65 70 75 80

Asp Val Lys Ser Val Lys Lys Glu Ile Trp Arg Gly Arg Arg Leu Lys 85 90 95

Cys Ser Phe Cys Asn Lys Gly Gly Ala Thr Val Gly Cys Asp Leu Trp 100 105 110

Phe Cys Lys Lys Ser Tyr His Tyr Val Cys Ala Lys Lys Asp Gln Ala 115 120 125

Ile Leu Gln Val Asp Gly Asn His Gly Thr Tyr Lys Leu Phe Cys Pro 130 135 140

Glu His Ser Pro Glu Gln Glu Glu Ala Thr Glu Ser Ala Asp Asp Pro 145 150 155 160

Ser Met Lys Lys Lys Arg Gly Lys Asn Lys Arg Leu Ser Ser Gly Pro 165 170 175

Pro Ala Gln Pro Lys Thr Met Lys Cys Ser Asn Ala Lys Arg His Met 180 185 190

Thr Glu Glu Pro His Gly His Thr Asp Ala Ala Val Lys Ser Pro Phe 195 200 205

Leu Lys Lys Cys Gln Glu Ala Gly Leu Leu Thr Glu Leu Phe Glu His 210 215 220

Ile Leu Glu Asn Met Asp Ser Val His Gly Arg Leu Val Asp Glu Thr 225 230 235 240

Ala Ser Glu Ser Asp Tyr Glu Gly Ile Glu Thr Leu Leu Phe Asp Cys 245 250 255

Gly Leu Phe Lys Asp Thr Leu Arg Lys Phe Gln Glu Val Ile Lys Ser 260 265 270

Lys Ala Cys Glu Trp Glu Glu Arg Gln Arg Gln Met Lys Gln Gln Leu 275 280 285

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Glu Asp His Gln

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<212> PRT

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<400> 46

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<213> Homo sapiens

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Lys	Met 50	Gln	Arg	Leu	Thr	Leu 55	Glu	His	Leu	Asn	Gln 60	Met	Val	Gly	Ile
Glu 65	Tyr	Ile	Leu	Leu	His 70	Ala	Gln	Glu	Pro	Ile 75	Leu	Phe	Ile	Ile	Arg 80
Lys	Gln	Gln	Arg	Gln 85	Ser	Pro	Ala	Ģln		Ile				Asp 95	
Tyr	Ile	Ile	Ala 100	Gly	Val	Ile		Gln 105	Ala	Pro	Asp		Gly 110	Ser	Val
Ile	Asn	Ser 115	Arg	Val	Leu	Thr	Ala 120	Val	His	Gly	Ile	Gln 125	Ser	Ala	Phe
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Lys	Arg	Lys		Glu 165	Pro	Ser	Ser	Ile	Phe 170	Gln	Arg	Gln	Arg	Val 175	
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Lys Ile Gly Gly Val Thr Glu Arg Met Pro Thr Pro Val Ile Lys Ala 85 90 95

Phe Gly Ile Leu Lys Arg Ala Ala Ala Glu Val Asn Gln Asp Tyr Gly 100 105 110

Leu Asp Pro Lys Ile Ala Asn Ala Ile Met Lys Ala Ala Asp Glu Val 115 120 125

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Gly Ser Gly Thr Gln Thr Asn Met Asn Val Asn Glu Val Ile Ser 145 150 155

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<212> PRT

<213> Homo sapiens

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Gln Lys Gln Pro Phe Ser Ile Glu Glu Ile Glu Val Ala Pro Pro Lys
35 40 45

Thr Lys Glu Val Arg Ile Lys Ile Leu Ala Thr Gly Ile Cys Arg Thr

Asp Asp His Val Ile Lys Gly Thr Met Val Ser Lys Phe Pro Val Ile 65 70 75 80

Val Gly His Glu Ala Thr Gly Ile Val Glu Ser Ile Gly Glu Gly Val

Thr Thr Val Lys Pro Gly Asp Lys Val Ile Pro Leu Phe Leu Pro Gln
100 105 110

Cys Arg Glu Cys Asn Ala Cys Arg Asn Pro Asp Gly Asn Leu Cys Ile 115 120 125

Arg Ser Asp Ile Thr Gly Arg Gly Val Leu Ala Asp Gly Thr Thr Arg 130 135 140

Phe Thr Cys Lys Gly Glu Pro Val His His Phe Met Asn Thr Ser Thr 145 150 155 160

Phe Thr Glu Tyr Thr

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<21	1> 2	59													
<21	2> P	RT													٠.
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Glu	Ser	Glu	Gln	Lys	Gly	Lys	Ala	Ala	Leu	Ala	Ala	Thr	Leu	Glu	Glu
1				- 5		_			10					15	
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															•
ıyr	гÀЗ	Ala	inr	vai	Ala	Ser	Asp		rre	GIU	met	ASI	-		ьуs
			20					25					30	•	
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Ala	Gln	T.eu	Glu				Gln		Val	Δla	Glu	Len	TVT	Ser	Tle
			014			2,0							_	-	
		35					40			•		45			
					45.1	i		- 13° r	,, :					_	
His	Asn	Ser	Gly	Asp	Lys	Ser	Asp	Ile	_Gln	`Asp	Leu	Leu	Glu	Ser	·Val
	਼.50					55					60				
	•						; .								115
B	T			~1									7	, ,	•
	rea	Asp	rys	GIU	_	AIA	Glu	THE	теп		Ser	Ser	ren	GIN	
65					70					75					80
		.:				4 4		.:		77.		1111		5 5 g/	1 14 .
Asp	Leu	Ala	His	Thr			Asp								
				85	5				90	3					
		•				_								95	
Ala	Lys	Val	Glu	Asp	Glu	Tyr	Arg	Ala	Phe	Gln	Glu	Glu	Ala	Lys	Lys
			100					105					.110		-
							5.4						:		
Gln	Tla						Thr				T.em				
GIII	116		wah.	, Leu	MSII	Mec		Tien	Giu	цуз	nen		Ser	, ASD	rieu
		115					120					125			_
				55.5		1 22	<u>.</u>	<u>:</u>	25.12			1.0	5 1	: .	
Asp	Glu	Lys	Glu				Ser								
	130					135		-		_	140				
*	~1		03	**- 1	~3	~1	***			17.1	T	T	*** -	•	
	GIU	Asp	GIU	vai		GIN	His	Arg			гÀ2	Leu	HIS	-	
145					150				٠.	155					160
												<i>.</i> .	7.	2	: '2'
Leu	Ile	Ile	Ser	Asp	Leu	Glu	Asn	Thr	Val	Lvs	Lvs	Leu	Gln		
				165					170	-4 -	-2-			175	
	***	1.					n ()								
Lys	His	Asp	Met	Glu.	Arg	Glu	Ile	Lys	Thr	Leu	His	Arg	Arg	Leu	Arg
			180					185					190		
												:			
~1	~1	0		~1		•								·	
GIU	GIU		ATA	GIU	пр		Gln	Pne	GII	ALA	Asp		GIH	m	Ald
		195	•				200					205			
	¥	:						: .	-, \			-		#	
Val	٧al	Tle	Δla	λen	Aen	Tle	Lys	Ser	Glu	Δla	Gln	Glin	Glu	Tle	Glv
·u1		116	ALG	veii	vəb		my 3	Ser	Giu	ALG		GIU	GIU	116	Gry
	210		•			215					220				
	5.5	7.	•		•			1			**				: •,*
Asp	Leu	Lys	Ara	Ara	Leu	His	Glu	Ala	Gln	Glu	Lvs	Asn	Glu	Lvs	Leu
225				- 3	230					235					240
					£ 3 V					درے					~4U
							٠.	٠.				. :		: .	: ~
Thr	Lys	Glu	Leu	Glu	Glu	Ile	Lys	Ser	Arg	Lys	Gln	Glu	Glu	Glu	Arg
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Gly Gly Tyr
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Thr Phe Tyr Arg Lys Glu Pro Phe Thr Leu Glu Ala Tyr Tyr Ser Ser 20 25 30

Pro Gln Asp Leu Pro Tyr Pro Asp Pro Ala Ile Ala Gln Phe Ser Val

Gln Lys Val Thr Pro Gln Ser Asp Gly Ser Ser Ser Lys Val Lys Val
50 55 60

Lys Val Arg Val Asn Val His Gly Ile Phe Ser Val Ser Ser Ala Ser 65 70 75 80

Leu Val Glu Val His Lys Ser Glu Glu Asn Glu Glu Pro Met Glu Thr 85 90 95

Asp Gln Asn Ala Lys Glu Glu Glu Lys Met Gln Val Asp Gln Glu Glu 100 105 110

Pro His Val Glu Glu Gln Gln Gln Gln Thr Pro Gly Arg 115 120 125

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<212> PRT

<213> Homo sapiens

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Tyr Val Val Cys Phe Ile Val Val Ala Gly Val Val Ile Leu Ala Val 20 25 30

Thr Ile Ala Leu Leu Val Tyr Phe Leu Ala Phe Asp Gln Lys Ser Tyr 35 40 45

Phe Tyr Arg Ser Ser Phe Gln Leu Leu Asn Val Glu Tyr Asn Ser Gln 50 55 60

Leu Asn Ser Pro Ala Thr Gln Glu Tyr Arg Thr Leu Ser Gly Arg Ile 65 70 75 80

Glu Ser Leu Ile Thr Lys Thr Phe Lys Glu Ser Asn Leu Arg Asn Gln
85 90 95

Phe Ile Arg Ala His Val Ala Lys Leu Arg Gln Asp Gly Ser Gly Val

Arg Ala Asp Val Val Met Lys Phe Gln Phe Thr Arg Asn Asn Gly
115 120 125

Ala Ser Met Lys Ser Arg Ile Glu Ser Val Leu Arg Gln Met Leu Asn 130 140

Asn Ser Gly Asn Leu Glu Ile Asn Pro Ser Thr Glu Ile Thr Ser Leu 145 155 160

Thr Asp Gln Ala Ala Ala Asn Trp Leu Ile Ash Glu Cys Gly Ala Gly 165 170 175

Pro Asp Leu Ile Thr Leu Ser Glu Gln Arg Ile Leu Gly Gly Thr Glu 180 185 190

Ala Glu Glu Gly Ser Trp Pro Trp Gln Val Ser Leu Arg Leu Asn Asn 195 200 205

Ala His His Cys Gly Gly Ser Leu Ile Asn Asn Met Trp Ile Leu Thr 210 220

Ala Ala His Cys Phe Arg Ser Asn Ser Asn Pro Arg Asp Trp Ile Ala 225 230 235 240

Thr Ser Gly Ile Ser Thr

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<211> 128

<212> PRT

<213> Homo sapiens

<400> 61

Gly Ile Phe Ser Val Ser Ser Ala Ser Leu Val Glu Val His Lys Ser 1 5 10 15

Glu Glu Asn Glu Glu Pro Met Glu Thr Asp Gln Asn Ala Lys Glu Glu 20 25

Glu Lys Met Gln Val Asp Gln Glu Glu Pro His Val Glu Glu Gln Gln 35 40 45

Gln Gln Thr Pro Ala Glu Asn Lys Ala Glu Ser Glu Glu Met Glu Thr 50 55 60

Ser Gln Ala Gly Ser Lys Asp Lys Lys Met Asp Gln Pro Pro Gln Ala 65 70 75 80

Lys Lys Ala Lys Val Lys Thr Ser Thr Val Asp Leu Pro Ile Glu Asn

90

95

Gln Leu Leu Trp Gln Ile Asp Arg Glu Met Leu Asn Leu Tyr Ile Glu 100 105 110

Asn Glu Gly Lys Met Ile Met Gln Asp Lys Leu Glu Lys Glu Arg Asn 115 120 125

<210> 62

<211> 418

<212> PRT

<213> Homo sapiens

<400> 62

Met Tyr Arg Pro Ala Arg Val Thr Ser Thr Ser Arg Phe Leu Asn Pro

Tyr Val Val Cys Phe Ile Val Val Ala Gly Val Val Ile Leu Ala Val 20 25 30

Thr Ile Ala Leu Leu Val Tyr Phe Leu Ala Phe Asp Gln Lys Ser Tyr
35 40 45

Phe Tyr Arg Ser Ser Phe Gln Leu Leu Asn Val Glu Tyr Asn Ser Gln 50 55 60

Leu Asn Ser Pro Ala Thr Gln Glu Tyr Arg Thr Leu Ser Gly Arg Ile
65 70 75 80

Glu Ser Leu Ile Thr Lys Thr Phe Lys Glu Ser Asn Leu Arg Asn Gln
85 90 95

Phe Ile Arg Ala His Val Ala Lys Leu Arg Gln Asp Gly Ser Gly Val

Arg Ala Asp Val Val Met Lys Phe Gln Phe Thr Arg Asn Asn Asn Gly

Ala Ser Met Lys Ser Arg Ile Glu Ser Val Leu Arg Gln Met Leu Asn 130 135 140

Asn Ser Gly Asn Leu Glu Ile Asn Pro Ser Thr Glu Ile Thr Ser Leu 145 150 155 160

Thr Asp Gln Ala Ala Ala Asn Trp Leu Ile Asn Glu Cys Gly Ala Gly
165 170 175

Pro Asp Leu Ile Thr Leu Ser Glu Gln Arg Ile Leu Gly Gly Thr Glu 180 185 190

Ala Glu Glu Gly Ser Trp Pro Trp Gln Val Ser Leu Arg Leu Asn Asn 195 200 205

Ala His His Cys Gly Gly Ser Leu Ile Asn Asn Met Trp Ile Leu Thr

210

215

220

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Ala 225	Ala	His	Cys		Arg 230	Ser	Asn	Ser	Asn	Pro 235	Arg	Asp	Trp	Ile	Ala 240	
Thr	Ser	Gly	Ile	Ser 245	Thr	Thr	Phe	Pro	Lys 250	Leu	Arg	Met	Arg	Val 255	_	
Asn	Ile	Leu	Ile 260	His	Asn	Asn	Tyr	Lys 265	Ser	Ala	Thr	His	Glu 270	Asn	Asp	
Ile	Ala	Leu 275	Val	Arg	Leu	Glu	Asn 280	Ser	Val	Thr	Phe	Thr 285	Lys	Asp	Ile	
His	Ser 290	Val	Cys	Leu	Pro	Ala 295	Ala	Thr	Gln		Ile 300	Pro	Pro	Gly	Ser	
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Gly 385	Ile	Val	Ser	Ţrp	Gly 390	Asp	Gln	Cys	Gly	Leu 395	Pro	Asp	Lys	Pro	Gly 400	
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aaca	gaaa	itt a	cago	agca	g cc	agca	acag	atg	gagg	ctc	aaga	taag	ag t	.cgca	aggaa	180
																240
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adta	LLGI	9	JULUC	.cgct	a ad	LLA	ıccyg	LCa	rygt	yee	aaag	guge	ya 9	julud	CCCCC	480

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 Ala Ser Gly Ser Leu Val Ala Thr Leu Gln Ser Leu Gly Ala Thr Gly
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Gly Asp Met Cys Lys Leu Lys Trp Val Glu Ile Ser Asn Glu Val Arg 65 70 75 80

Thr Glu Ser His Met Asp Trp Glu Lys Val Ala Phe Lys Asp Phe Ser

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	Lys	Asn	Pro	Туг 100	Lys	Gly	Lys	Lys	Leu 105	Lys	Lys	His	Pro	Asp 110	Phe	Pro
	Lys	Lys	Pro 115		Thr	Pro	Tyr	Phe 120	Arg	Phe	Phé	Met	Glu 125	Lys	Arg	Ala
ť.	Lys	Tyr 130		Lys	Leu	His	Pro 135	Glu	Met	Ser	Asn	Leu 140	Asp	Leu	Thr	Lys
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	Ser	Asp	Ile 195	Pro	Glu	Lys	Pro	Lys 200	Thr	Pro	Gln	Gln	Leu 205	Trp	Tyr	Thr
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				260					265					Leu 270		ţ
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	•	•.	35 5					360				٠.	365	Pro		
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Ly	s Gl	y Gl	y Se	r Gl 40		s Pro	o Ly:	s Ar	g Pro		l Se	r Al	a Me	Ph 41	
Phe	e Se	r Gl	u Gl: 42	u Ly: 0 :::	s Arg	g Arg	g Glr			n Gl	u Gli	ı Ar	g Pro 430		u Lei
Ser	Gl:	1 Se 43	r Gl	u Le	u Thi	Arg	140	Lei	u Ala	Ar	g Met	Tr ₁	p Asr 5) Le
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Ala 465	Glr i	ı Se	r Glu	ı Arg	470	Pro	Gly	Gly	- / Glu	475		Glu	ı Arg	Gly	Lys 480
Leu	Pro	Gl	u Sei	485	Lys i	Arg	Ala	Glu	490		e Trp	Gln	ı Gln		(*. :/***
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Lys	Ala	Me1	Glu 5	Met	Thr	Trp	Asn 520	Asn	Met				Glu		
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Met	Lys	Phe	Gln	Gly 565	Glu	Pro	ГÁЗ	Lys	Pro 570	Pro	Met	Asn	Gly	Tyr 575	
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Asp	Arg	Ala	Ala	Tyr 645	Lys	Glu	Tyr	Ile	Ser 650	Asn	Lys	Arg	Lys		
Thr	Lys	Leu	Arg 660	Gly	Pro	Asn		Lys 665	Ser	Ser	Arg	Thr	Thr 670	Leu	Gln

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Trp Pro Pro Gly Arg Ser Leu Pro Lys Ala Val Arg Pro Pro Leu Ser

Trp Pro Pro Phe Ser Gln Gln Gln Thr Leu Pro Val Met Ser Gly Glu 105

100

90

35.

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Leu Ser Cys Glu Arg Leu Arg Ala Leu Leu Pro Gln Phe Asp Gly Arg

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75

70

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Secretary Secretary Secretary
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(P) 65 기 (대한 제한 ) 전 4 70 인간 (대한 ) 전 175 기 (대한 ) 4 (대한 80)
ent for the second
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  Lys Pro Gly Asp Lys Val Ile Pro Leu Phe Leu Pro Gln Cys Arg Glu
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Thr 385	Phe			.74	, .	•					- 20	* * •		`e	. · ·	
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		Ile			e Vertiger	: : :					•		-,				
	<21	0>, 8.	ġ.,		pr ¹ ·		÷					· ·	٠,			•	
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Asn Asn Leu Glu Phe Ala Lys Glu Leu Gln Arg Ser Phe Met Ala Leu
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Ser Gln Asp Ile Gln Lys Thr Ile Lys Lys Thr Ala Arg Arg Glu Gln
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Leu Met Arg Glu Glu Ala Glu Gln Lys Arg Leu Lys Thr Val Leu Glu
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Leu Gln Tyr Val Leu Asp Lys Leu Gly Asp Asp Glu Val Arg Thr Asp
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Leu Pro Gly Val Val Ile Leu Ala Val Pro Ile Ala Leu Leu Val Tyr 50 55 60

Phe Leu Ala Phe Asp Gln Lys Ser Tyr Phe Tyr Trp Ser Asn Phe Pro 65 70 75 80

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Pro Val Gly Glu Asp Ser Leu Val Ser Asp Arg Ser Lys Lys Glu Leu 35 40 45

Ser Pro Val Leu Thr Ser Glu Val His Ser Val Arg Ala Gly Arg His
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Leu Ala Thr Lys Leu Asn Ile Leu Val Gln Gln His Phe Asp Leu Ala 65 70 75 80

Ser Thr Thr Ile Thr Asn Ile Pro Met Lys Glu Glu Gln His Ala Asn 85 90 95

Thr Ser Ala Asn Tyr Asp Val Glu Leu Leu His His Lys Asp Ala His 100 105 110

Val Asp Phe Leu Lys Ser Gly Asp Ser His Leu Gly Gly Gly Ser Arg 115 120 125

Glu Gly Ser Phe Lys Glu Thr Ile Thr Leu Lys Trp Cys Thr Pro Arg 130 135 140

Thr Asn Asn Ile Glu Leu His Tyr Cys Thr Gly Ala Tyr Arg Ile Ser 145 150 155 160

Pro Val Asp Val Asn Ser Arg Pro Ser Ser Cys Leu Thr Asn Phe Leu 165 170 175 Leu Asn Gly Arg Ser Val Leu Leu Glu Gln Pro Arg Lys Ser Gly Ser 180 185 190

Lys Val Ile Ser His Met Leu Ser Ser His Gly Gly Glu Ile Phe Leu 195 200 205

His Val Leu Ser Ser Ser Arg Ser Ile Leu Glu Asp Pro Pro Ser Ile 210 215 220

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35 40 45

Asp Asn Asn His Leu Ile Gln Cys Ile Met Asp Ser Gln Asn Lys Gly
50 55 60

Lys Thr Ser Glu Cys Ser Gln Tyr Gln Gln Met Leu His Thr Asn Leu 65 70 75 80

Val Tyr Leu Ala Thr Ile Ala Asp Ser Asn Gln Asn Met Gln Ser Leu 85 90 95

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Glu Lys Ile Ser Gln Pro Val Val Val Val Ala Ile Val Gly Leu Tyr
50 55 60

Arg Thr Gly Lys Ser Tyr Leu Met Asn His Leu Ala Gly Gln Asn His 65 70 75 80

Gly Phe Pro Leu Gly Ser Thr Val Gln Ser Glu Thr Lys Gly Ile Trp 85 90 95

Met Trp Cys Val Pro His Pro Ser Lys Pro Asn His Thr Leu Val Leu 100 105 110

Leu Asp Thr Glu Gly Leu Gly Asp Val Glu Lys Gly Asp Pro Lys Asn 115 120 125

Asp Ser Trp Ile Phe Ala Leu Ala Val Leu Leu Cys Ser Thr Phe Val 130 135 140

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Lys Ile Ser Gln Pro Val Val Val Val Ala Ile Val Gly Leu Tyr Arg
35 40 45

Thr Gly Lys Ser Tyr Leu Met Asn His Leu Ala Gly Gln Asn His Gly 50 55 60

Phe Pro Leu Gly Ser Thr Val Gln Ser Glu Thr Lys Gly Ile Trp Met 65 70 75 80

Trp Cys Val Pro His Pro Ser Lys Pro Asn His Thr Leu Val Leu Leu

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Asp Thr Glu Gly Leu Gly Asp Val Glu Lys Gly Asp Pro Lys Asn Asp 100 105 110

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Ile Asn Asp Pro Phe Ile Asp Leu Asn Tyr Met Val Tyr Met Phe Gln
35 40 45

Tyr Asp Ser Thr His Gly Lys Phe His Gly Thr Val Glu Ala Glu Asn

Gly Lys Leu Val Ile Asn Gly Asn Pro Ile Thr Ile Phe Gln Glu Arg
65 70 75 80

Asp Pro Ser Lys Ile Lys Trp Gly Asp Ala Gly Ala Glu Tyr Val Val 85 90 95

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                         55
                                              60
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Glu Asn Gly Lys Leu Val Ile Asn Gly Asn Pro Ile Thr Ile Phe Gln
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<213> Homo sapiens

<400> 112

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Gln	Gl u	Phe 35		Trp	Asp	Tyr	Val		Leu	Asp	Glu	Ala 45		Lys	Ile
Lys	Thr 50		Ser	Thr	Lys	Ser 55		Ile	Cys	Ala	Arg 60		Ile	Pro	Ala
	Asn 5	Arg	Leu	Leu	Leu 70		Gly	Thr	Pro	Ile 75	Gln	Asn	Asn	Leu	Gln 80
Glu	Leu	Trp	Ser	Leu 85	Phe		Phe	Ala	Cys 90		Gly	Ser	Leu	Leu 95	_
Thr	Leu	Lys	Thr 100	Phe	Lys	Met	Glu	Tyr 105		Asn	Pro	Ile	Thr 110	Arg	Ala
Arg	Glu	Lys 115		Ala	Thr	Pro	Gly 120		Lys	Ala	Leu	Gly 125	Phe	Lys	Ile
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Lys 145		Asp		Gln	Lys 150				Ser		Pro	Glu	Ala		Leu 160
Asn	Glu		Asn		Asp ///					Cys		Met	Pro	Ser 175	
Ser		Arg			Leu						Leu		Pro 190		Gln
Glu		Ile 195		Arg	Lys · *:			Ser		Asp		Ile 205	Lys	Glu	Leu
Leu					Ser							Val	Leu	Lys	Lys
225	सन्द	25 81. 15		• •	Arg 230	53	Leu	Ser		Arg 235	Ala	Суз	Cys		Leu 240
Asn	Leu	Gly		Phe 245	Ser	Ala			٠.				5 (12) P (13)		

<212> PRT

<213> Homo sapiens

<400> 113

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Thr	Tyr	Ile 35	Pro	Lys	Asp	Ser	Lys 40	Lys	Lys	Lys	His	Glu 45	Leu	Lys	Ile
Thr	Gln 50	Gln	Gly	Thr	Asp	Pro 55	Leu	Val	Leu	Ala	Val 60	Gln	Ser	Lys	Glu
Gln 65	Ala	Glu	Gln	Trp	Leu 70		Val	Ile	Lys	Glu 75	Ala	Tyr	Ser	Gly	Cys 80
Ser	Gly	Pro	Val	Asp 85	Ser	Glu	Суз	Pro	Pro 90	Pro	Pro	Ser	Ser	Pro 95	Va]
His	Lys	Ala	Glu 100	Leu	Glu	Lys	Lys	Leu 105		.Ser				•	
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Leu	Gly	Asp 35	Asp	Ile	Asp	Leu	Ile 40	Val			Glu	His 45	Asp	Gly	Va]
								, ,	: 					Min en	T
Met	Thr 50	Gly	Ala	Asn	Gly	55	Val	Ser	Pne	He	Asn 60	11e	гÀз	inr	rec
Asn 65		Trp	Asp	Ser	Arg 70	His	Суз	neA	Gly	Va1 75		Trp	Arg	Gln	Lys 80
Leu	Asp	Ser	Gln	Arg 85		Ala	Val	Ile	Ala · 90			Leu		95	
Ser	Tyr	Lys	Leu 100	Ala	Arg	Trp	Thr	Cys 105	Cys	Ala	Leu				
Glu	Tyr	Leu 115	Lys	Leu	Gly	Tyr	Val 120	Ser	Arg	Tyr	His	Val 125	Lys	Asp	Sei
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cetetgeage tgetggaege etectggtae etgeegaage tggggegega egegegaege 180
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 Asp Leu Pro Ala Ser Thr Pro Ala Ser Lys Ser Cys Asp Ser Ser Pro
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 Pro Gln Asp Ala Ser Thr Pro Arg Pro Ser Ser Ala Ser His Leu Cys
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Gln Leu Ala Ala Lys Pro Ala Pro Ser Thr Asp Ser Val Ala Leu Arg
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Ser Pro Leu Thr Leu Ser Ser Pro Phe Thr Thr Ser Phe Ser Leu Gly
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                                105
                                                    110
Ser His Ser Thr Leu Asn Gly Asp Leu Ser Val Pro Ser Ser Tyr Val
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Ser Leu His Leu Ser Pro Gln Val Ser Ser Ser Val Val Tyr Gly Arg
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Ser Pro Val Met Ala Phe Glu Ser His Pro His Leu Arg Gly Ser Ser
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Asp Gly Lys Val

Asp Gly Lys Val

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35 40

Phe Glu Glu Arg His Ile Pro Gly Ala Ala Phe Phe Asp Ile Asp Gln 50 60

Cys Ser Asp Arg Thr Ser Pro Tyr Asp His Met Leu Pro Gly Ala Glu 65 70 75 80

His Phe Ala Glu Tyr Ala Gly Arg Leu Gly Val Gly Ala Ala Thr His 85 90 95

Val Val Ile Tyr Asp Ala Ser Asp Gln Gly Leu Tyr Sèr Ala Pro Arg 100 105 110

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Gly Ala Pro His Asn Pro Ala Pro Pro Thr Ser Thr Val Ile His Ile 35 40 45

A	rg S	er G 50	lu T	hr s	er V	al I	Pro 55	Asp	Hi	s Va	al V	al '	Trp 60		ŕ L	eu É	he	Asn	.,
TI	nr L	eu P	he M	et A	sn P	ro (Cys	Cys	Le	u ·G]		he 3	Πe	Al	a Pi	ne A	la	Tyr 80	
Se	er Va	al Ly	ys S	er A	rg A 85	sp A	arg	Lys	Me		1 G 0	ly I	\sp	V a	l Ti		ly 95	Ala	
G1	n Al	ia Ty	/r A	la Se DO	er T	hr A	la	Lys	Cy:	s Le 5	u As	sn I	le	Tr		a L	eu	Ile	
Le	u Gl	y II 11	.e Le .5	eu Me	t T	hr I	le	Leu 120	Le	ıIl	e Vá	al I		Pro		1 L	eù	Ile	
Ph	e Gl	n Al	а Тъ	cr G3	3.5						. •								
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Arg	Île	Leu	Thr	Glu	Leu	ı Th	r T	hr 1	Mr	λla	The	∷ • Th	 ∽ •	h.	7 T -	».	and and	ħr	۔
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Thr 145		Met	Ala					•			٠ .				٠.		***	er	

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304°

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345

030

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1.3.5

1.4

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<213> homo sapien

<400> 158

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                                                                    300
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                                                                    360
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                                                                    420
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                                                                    720
cctagagete taettacaga acageeetga ggeetgtgae tatgggetet gaagggggea
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gegtgataag gagaaggagg tggaatgtca geaggageat atceatgaae teeaggaget
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                                                                      360
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                                                                      180
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                                                                      240
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agtgaagaag ttccagcttt tggggtagca tctccgcccc cacttactga tactcctgac
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                                                                      360
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                                                                     120
                                                                     180
agatggtgaa agaaacaact tactacgatg ttttgggggt caaacccaat gctactcagg
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aaagggaatt atatgacaaa ggaggagaac aggcaattaa agagggtgga gcaggtggcg
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gttttggctc ccccatggac atctttgata tgttttttgg aggaggagga aggatgcaga
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893
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      <211> 549
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                                                7 100
      <213> homo sapien
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                                                                     240
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ttctatgggt caaaagagga tccacagact ttctattatg ctgttgctgt ggtgaagaag
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aaacctc ...
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                                                                    540
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ggaagatccc acagtctcag ccctgcttac tagtgagaaa gactggcaag gtttcctaga
                                                                    720
                                                                    780
gctctactta cagaacagcc ctgaggcctg tgactatggg ctctgaaggg ggcaggagtc
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-

838

14. B.

1

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3563

254

V8

 $(W)^{n}$

130

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                                                                         240
  ctacaacate cagaaagagt ccaccetgca cetggtgete egteteagag gtgggatgea
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                                                                         360
categagaac gteaaageaa agateeagga caaggaagge attecteetg accageagag
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493

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Lys Met Glu Glu Glu Ser Gly Ala Pro Cys Val Pro Ser Gly Asn Gly Ala Pro Gly Pro Lys Gly Glu Glu Arg Pro Thr Gln Asn Glu Lys Arg 40 45 Lys Glu Lys Asn Ile Lys Arg Gly Gly Asn Arg Phe Glu Pro Tyr Ser - 60 Asn Pro Thr Lys Arg Tyr Arg Ala Phe Ile Thr Asn Ile Pro Phe Asp 70 Val Lys Trp Gln Ser Leu Lys Asp Leu Val Lys Glu Lys Val Gly Glu 85 90 Val Thr Tyr Val Glu Leu Leu Met Asp Ala Glu Gly Lys Ser Arg Gly 100 105 Cys Ala Val Val Glu Phe Lys Met Glu Glu Ser Met Lys Lys Ala Ala 120 125 Glu Val Leu Asn Lys His Ser Leu Ser Gly Arg Pro Leu Lys Val Lys \$ 1.50 PM 135 140 Glu Asp Pro Asp Gly Glu His Ala Arg Arg Ala Met Gln Lys Ala Gly --150 155 Arg Leu Gly Ser Thr Val Phe Val Ala Asn Leu Asp Tyr Lys Val Gly 165 170 175 Trp Lys Lys Leu Lys Glu Val Phe Ser Met Ala Gly Val Val Val Arg *→* 185 - 180 190 Ala Asp Ile Leu Glu Asp Lys Asp Gly Lys Ser Arg Gly Ile Gly Ile 195 200 Val Thr Phe Glu Gln Ser Ile Glu Ala Val Gln Ala Ile Ser Met Phe 210 215 220 Asn Gly Gln Leu Leu Phe Asp Arg Pro Met His Val Lys Met Asp Glu 230 235 Arg Ala Leu Pro Lys Gly Asp Phe Phe Pro Pro Glu Arg His Ser 245 250

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Lys Thr Glu Ser Val Lys Glu Ser Glu Ser Leu Met Glu Phe Ala Gln 50 55 60
Pro Glu Ile Gln Pro Gln Glu Phe Leu Asn Arg Arg Tyr Met Thr Glu

65 70 75 80

Val Asp Tyr Ser Asn Lys Gln Gly Glu Glu Gln Pro Trp Glu Ala Asp
85 90 95

Tyr Ala Arg Lys Pro Asn Leu Pro Lys Arg Trp Asp Met Leu Thr Glu 100 105 110

Pro Asp Gly Gln Glu Lys Lys Gln Glu Ser Phe Lys Ser Trp Glu Ala 115 120 125

Ser Gly Lys His Gln Glu Val Ser Lys Pro Ala Val Ser Leu Glu Gln
130
135
140
Arg Lys Gln Asp The Ser Lys Low And Ser The Lys Rep Glo Gl

Arg Lys Gln Asp Thr Ser Lys Leu Arg Ser Thr Leu Pro Glu Glu Gln
145 150 155 160

Lys Lys Gln Glu Ile Ser Lys Ser Lys Pro Ser Pro Ser Gln Tro Lys
165 170 175

Gln Asp Thr Pro Lys Ser Lys Ala Gly Tyr Val Gln Glu Glu Gln Lys 180 185 190

Lys Gln Glu Thr Pro Lys Leu Trp Pro Val Gln Leu Gln Lys Glu Gln
195 200 205

Asp Pro Lys Lys Gln Thr Pro Lys Ser Trp Thr Pro Ser Met Gln Ser 210 215 220

Glu Gln Asn Thr Thr Lys Ser Trp Thr Thr Pro Met Cys Glu Glu Gln 225 230 235 240

Asp Ser Lys Gln Pro Glu Thr Pro Lys Ser Trp Glu Asn Asn Val Glu 245 250 255

Ser Gln Lys His Ser Leu Thr Ser Gln Ser Gln Ile Ser Pro Lys Ser 260 265 270

Trp Gly Val Ala Thr Ala Ser Leu Ile Pro Asn Asp Gln Leu Leu Pro 275 280 285

Arg Lys Leu Asn Thr Glu Pro Lys Asp Val Pro Lys Pro Val His Gln 290 295 300

Pro Val Gly Ser Ser Ser Thr Leu Pro Lys Asp Pro Val Leu Arg Lys 305 310 315 320

Glu Lys Leu Gln Asp Leu Met Thr Gln Ile Gln Gly Thr Cys Asn Phe 325 330 335

Met Gln Glu Ser Val Leu Asp Phe Asp Lys Pro Ser Ser Ala Ile Pro Thr Ser Gln Pro Pro Ser Ala Thr Pro Gly Ser Pro Val Ala Ser Lys Glu Gln Asn Leu Ser Ser Gln Ser Asp Phe Leu Gln Glu Pro Leu Gln Val Phe Asn Val Asn Ala Pro Leu Pro Pro Arg Lys Glu Gln Glu Ile Lys Glu Ser Pro Tyr Ser Pro Gly Tyr Asn Gln Ser Phe Thr Thr Ala Ser Thr Gln Thr Pro Pro Gln Cys Gln Leu Pro Ser Ile His Val Glu Gln Thr Val His Ser Gln Glu Thr Ala Ala Asn Tyr His Pro Asp Gly Thr Ile Gln Val Ser Asn Gly Ser Leu Ala Phe Tyr Pro Ala Gln Thr Asn Val Phe Pro Arg Pro Thr Gln Pro Phe Val Asn Ser Arg Gly Ser Val Arg Gly Cys Thr Arg Gly Gly Arg Leu Ile Thr Asn Ser Tyr Arg Ser Pro Gly Gly Tyr Lys Gly Phe Asp Thr Tyr Arg Gly Leu Pro Ser Ile Ser Asn Gly Asn Tyr Ser Gln Leu Gln Phe Gln Ala Arg Glu Tyr Ser Gly Ala Pro Tyr Ser Gln Arg Asp Asn Phe Gln Gln Cys Tyr Lys 535 ... Arg Gly Gly Thr Ser Gly Gly Pro Arg Ala Asn Ser Arg Ala Gly Trp Ser Asp Ser Ser Gln Val Ser Ser Pro Glu Arg Asp Asn Glu Thr Phe Asn Ser Gly Asp Ser Gly Gln Gly Asp Ser Arg Ser Met Thr Pro Val Asp Val Pro Val Thr Asn Pro Ala Ala Thr Ile Leu Pro Val His Val Tyr Pro Leu Pro Gln Gln Met Arg Val Ala Phe Ser Ala Ala Arg Thr Ser Asn Leu Ala Pro Gly Thr Leu Asp Gln Pro Ile Val Phe Asp Leu Leu Leu Asn Asn Leu Gly Glu Thr Phe Asp Leu Gln Leu Gly Arg Phe Asn Cys Pro Val Asn Gly Thr Tyr Val Phe Ile Phe His Met Leu Lys Leu Ala Val Asn Val Pro Leu Tyr Val Asn Leu Met Lys Asn Glu Glu Val Leu Val Ser Ala Tyr Ala Asn Asp Gly Ala Pro Asp His Glu Thr Ala Ser Asn His Ala Ile Leu Gln Leu Phe Gln Gly Asp Gln Ile Trp Leu Arg Leu His Arg Gly Ala Ile Tyr Gly Ser Ser Trp Lys Tyr Ser Thr Phe Ser Gly Tyr Leu Leu Tyr Gln Asp

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GLY.	Gia	vah	100	a.y	014		0	105		•			110	,	
C	D	3		T	Co	0350	Glu		Dro	Cure	Thr	G) n		Ser	Glv
Ser.	PIO		ASD	ren	Ser	GIY		Ser	FIO	Cys	1111	125	мy	DCT	GIY
_ `		115				-3	120	0	<u>.</u>		n		Minima.	Dia	C
Leu		Pro	Glu	Arg	Arg		Asp	Ser	Pro	irp		Pro	11p	PIO	Ser
2.7	130					135	_	_ ,	_		140		417		
Pro	Gln	Glu	Arg	Asp			Thr	Arg	Asp		GIU	GIU	Ser	PTO	
145	٠.				150	, 3€.				155		23			160
Asp	Trp	Gly	Gly	Ala	Glu	Ser	Pro	Arg	Gly	Trp	Glu	Ala	Gly	Pro	Arg
				165			۲ کی در د		170		•			175	
Glu	Trp	Gly	Pro	Ser	Pro	Ser	Gly	His	Gly	Asp	Gly	Pro	Arg	Arg	Arģ
•			180					185					190	4 - 7	57
Pro	Arg	Lys	Arg	Arg	Gly	Arg	Lys	Gly	Arg	Met	Gĺy	Arg	Gln	His	Glu
		195	-			•	200		***	· •		205			٠.
Ala	Ala	Ala	Thr	Ala	Ala	Thr	Ala	Ala	Thr	Ala	Thr	Gly	Gly	Thr	Ala
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	Glv	Ara	Ala	Arai	Glv	Pro	Arg	Gln	Gln	Glý	Arq	Arg	Arq	His	Gly
		3		245			ું કે	e* **	250	•				255	
Thr	Gln	Ara	Ara		Glv	Pro	Pro	Gln		Arg	Glu	Glu	Glv	Pro	Arg
	V	3	260		,			265		3			270		
Agn	αla	Thr		Tle	Leu	Glv	Leu		Thr	Pro	Ser	Glv	Glu	Gln	Arq
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A1 =) den		Ser	Gln	Δla	Leu	Pro	Ala	Leu	Ala	Glv	,	Ala	Ala	Ala
	290		001	0111		295					300				
		uia	a [4	714	Pro		Ala	Ġlv	Pro	Alá		Ala	Pro	Val	Glv
305	ALU		· ALG	-	310	017	****	017		315	,				320.
			2	2-0		Clv	Trp	7	Gly	-	Ara	Arm	ČIÜ	ัดเจ	
GIA	ALG	GIY					LLP	<u>.</u>	330	OLY.	****			335	-,
		. 3 3 0					Arg	· C1		A~~	Gly	A*~	ัดโซ	-	Glv
AId	GIY.	ALG			Gry	GIY	MY	345	GLY	MLG	GIA	ALY.	350	فسف	U-y
~ 3	~ 3		340		01	@1.4			X	61.	C1	63.5		A1 a	GIV
GIA.	GIY.	-	GIY	GIY	GIĀ	GIY	Ala	GIÅ	Arg	GLY	GIŸ		WTG	Mra.	GLY
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Pro		Glu	Gly	ALa	Ser			GIY	ALS	arg		GIY	GTĤ	GIII	Arg
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385					390					395			<u> </u>	الأعالى إ	400
Arg	Gly	Arg	Arg	Ala	Arg	Gly	Gln	Arg	Ala	Gly	Glu	Glu	Ala		Asp
				405					410			۰۰۰ مرود پا		415	
Gly	Leu	Leu	Pro	Arg	Gly	Arg	Asp	Arg	Leu	Pro	Leu	Arg	Pro	Gly	Asp
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Ala	Asn	Gln	Arg	Ala	Glu	Arg	Pro	Gly	Pro	Pro	Arg	Gly	Gly	His	Gly
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465					470			٠.		475	, -				480
	Glv	Glv	Glv	Phe			Pro	Pro			Ara	Pro	Pro	Ala	Val
	1	2	1	485					490		3			495	
Tæn	T.e.	Pro	T.e.v		Arcr	Len	Ala	Cva			Asp	Pro	Glv	-	Thr
 u			500		3			505		,			510		
			200					553							

Arg Pro Gly Pro Arg Arg Pro Ala Arg Arg Pro Arg Gly Glu Leu Ile 515 520 525 Pro Arg Arg Pro Asp Pro Ala Ala Pro Ser Glu Glu Gly Leu Arg Met 535 540 Glu Ser Ser Val Asp Asp Gly Ala Thr Ala Thr Thr Ala Asp Ala Ala 550 555 Ser Gly Glu Ala Pro Glu Ala Gly Pro Ser Pro Ser His Ser Pro Thr 565 570 Met Cys Gln Thr Gly Gly Pro Gly Pro Pro Pro Pro Gln Pro Pro Arg 585 590 Trp Leu Pro 595

<210> 188 <211> 376 <212> PRT <213> Homo sapien

<400> 188 Glu Met Arg Lys Phe Asp Val Pro Ser Met Glu Ser Thr Leu Asn Gln 1 10 15 Pro Ala Met Leu Glu Thr Leu Tyr Ser Asp Pro His Tyr Arg Ala His 20 1. The second of the control of the second o Phe Pro Asn Pro Arg Pro Asp Thr Asn Lys Asp Val Tyr Lys Val Leu 40 45 45 45 Pro Glu Ser Lys Lys Ala Pro Gly Ser Gly Ala Val Phe Glu Arg Asn 55 60 Gly Pro His Ala Ser Ser Ser Gly Val Leu Pro Leu Gly Leu Gln Pro . 70 Ala Pro Glý Leu Ser Lys Ser Leu Ser Ser Gln Val Trp Gln Pro Ser g 900 grant 195 c. Pro Asp Pro Trp His Pro Gly Glu Gln Ser Cys Glu Leu Ser Thr Cys 105 100 110 Arg Gln Gln Leu Glu Leu Ile Arg Leu Gln Met Glu Gln Met Gln Leu 120 1.50 get 200 (125 to 10 10 to Gln Asn Gly Ala Met Cys His His Pro Ala Ala Phe Ala Pro Leu Leu Pro Thr Leu Glu Pro Ala Gln Trp Leu Ser Ile Leu Asn Ser Asn Glu 150 155 His Leu Leu Lys Glu Leu Leu Ile Asp Lys Gln Arg Lys His 165 170 **175** Ile Ser Gln Leu Glu Gln Lys Val Arg Glu Ser Glu Leu Gln Val His 180 185 190 L Ser Ala Leu Leu Gly Arg Pro Ala Pro Phe Gly Asp Val Cys Leu Leu 200 205 Arg Leu Gln Glu Leu Gln Arg Glu Asn Thr Phe Leu Arg Ala Gln Phe 210 215 220 Ala Gln Lys Thr Glu Ala Leu Ser Lys Glu Lys Met Glu Leu Glu Lys 230 235 Lys Leu Ser Ala Ser Glu Val Glu Ile Gln Leu Ile Arg Glu Ser Leu 245 250 Lys Val Thr Leu Gln Lys His Ser Glu Glu Gly Lys Lys Gln Glu Glu 265 Arg Val Lys Gly Arg Asp Lys His Ile Asn Asn Leu Lys Lys Lys Cys

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Gln Lys Glu Ser Glu Gln Asn Arg Glu Lys Gln Gln Arg Ile Glu Thr
                      295
                                        300
Leu Glu Arg Tyr Leu Ala Asp Leu Pro Thr Leu Glu Asp His Gln Lys
               310
                                   315
Gln Thr Glu Gln Leu Lys Asp Ala Glu Leu Lys Asn Thr Glu Leu Gln
          325
                                330
Glu Arg Val Ala Glu Leu Glu Thr Leu Leu Glu Asp Thr Gln Ala Thr
           340
                         345
                                   350
Cys Arg Glu Lys Glu Val Gln Leu Glu Ser Leu Arg Gln Arg Glu Ala
 . . . . . 355
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Asp Leu Ser Ser Ala Arg His Arg
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                      375
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<210> 189 <211> 160 <212> PRT <213> Homo sapien

<400> 189

Met Leu Glu Ala His Arg Arg Gln Arg His Pro Phe Leu Leu Gly 1 Thr Thr Ala Asn Arg Thr Gln Ser Leu Asn Tyr Gly Cys Ile Val Glu 30 Asn Pro Gln Thr His Glu Val Leu His Tyr Val Glu Lys Pro Ser Thr 9 10 35 35 3 1 1 1 4 9 1 40 4 1 5 1 1 1 1 45 Phe Ile Ser Asp Ile Ile Asn Cys Gly Ile Tyr Leu Phe Ser Pro Glu 50 m 20 m 60 m Ala Leu Lys Pro Leu Arg Asp Val Phe Gln Arg Asn Gln Gln Asp Gly 75 Gln Leu Glu Asp Ser Pro Gly Leu Trp Pro Gly Ala Gly Thr Ile Arg 90. Leu Glu Gln Asp Val Phe Ser Ala Leu Ala Gly Gln Gly Gln Ile Tyr. 100. Val His Leu Thr Asp Gly Ile Trp Ser Gln Ile Lys Ser Ala Gly Ser 120 Ala Leu Tyr Ala Ser Arg Leu Tyr Leu Ser Arg Tyr Gln Asp Thr His Pro Glu Arg Leu Ala Lys His Thr Pro Gly Gly Pro Trp Ile Arg Gly 145 150 155

<210> 190 <211> 146 <212> PRT <213> Homo sapien

<400> 190

 Met Asp Pro Arg Ala Ser Leu Leu Leu Leu Gly Asn Val Tyr Île His

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 Pro Thr Ala Lys Val Ala Pro Ser Ala Val Leu Gly Pro Asn Val Ser 20
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 30

 Ile Gly Lys Gly Val Thr Val Gly Glu Gly Val Arg Leu Arg Glu Ser 35
 40
 45

 Ile Val Leu His Gly Ala Thr Leu Gln Glu His Thr Cys Val Leu His 50
 55
 60

 Ser Ile Val Gly Trp Gly Ser Thr Val Gly Arg Trp Ala Arg Val Glu

75 Gly Thr Pro Ser Asp Pro Asn Pro Asn Asp Pro Arg Ala Arg Met Asp -85 90 Ser Glu Ser Leu Phe Lys Asp Gly Lys Leu Leu Pro Ala Ile Thr Ile 100 105 . . Leu Gly Cys Arg Val Arg Ile Pro Ala Glu Val Leu Ile Leu Asn Ser 115 120 125 Ile Val Leu Pro His Lys Glu Leu Ser Arg Ser Phe Thr Asn Gln Ile 130 135 Ile Leu 145 Jajanes

<210> 191 <211> 704 <212> PRT <213> Homo sapien

<400> 191 Glu Gly Gly Cys Ala Ala Gly Arg Gly Arg Glu Leu Glu Pro Glu Leu 10 15 Glu Pro Gly Pro Gly Pro Gly Ser Ala Leu Glu Pro Gly Glu Glu Phe 20 Glu Ile Val Asp Arg Ser Gln Leu Pro Gly Pro Gly Asp Leu Arg Ser 35 40 1 45 45 W Ala Thr Arg Pro Arg Ala Ala Glu Gly Trp Ser Ala Pro Ile Leu Thr j - 1 j **60** j j - 1 m - 1 m Leu Ala Arg Arg Ala Thr Gly Asn Leu Ser Ala Ser Cys Gly Ser Ala 70 75 Leu Arg Ala Ala Gly Leu Gly Gly Gly Asp Ser Gly Asp Gly Thr 95 90 Ala Arg Ala Ala Ser Lys Cys Gln Met Met Glu Glu Arg Ala Asn Leu Met His Met Met Lys Leu Ser Ile Lys Val Leu Leu Gln Ser Ala Leu .. 115 120 Ser Leu Gly Arg Ser Leu Asp Ala Asp His Ala Pro Leu Gln Gln Phe 135 . . . 140 Phe Val Val Met Glu His Cys Leu Lys His Gly Leu Lys Val Lys Lys 150 155 Ser Phe Ile Gly Gln Asn Lys Ser Phe Phe Gly Pro Leu Glu Leu Val 165 170 Glu Lys Leu Cys Pro Glu Ala Ser Asp Ile Ala Thr Ser Val Arg Asn 180 185 190 Leu Pro Glu Leu Lys Thr Ala Val Gly Arg Gly Arg Ala Trp Leu Tyr 200 205 Leu Ala Leu Met Gln Lys Lys Leu Ala Asp Tyr Leu Lys Val Leu Ile 215 220 Asp Asn Lys His Leu Leu Ser Glu Phe Tyr Glu Pro Glu Ala Leu Met 230 235 240 Met Glu Glu Gly Met Val Ile Val Gly Leu Leu Val Gly Leu Asn 245 250 Val Leu Asp Ala Asn Leu Cys Leu Lys Gly Glu Asp Leu Asp Ser Gln 260 265 Val Gly Val Ile Asp Phe Ser Leu Tyr Leu Lys Asp Val Gln Asp Leu 280 285 Asp Gly Gly Lys Glu His Glu Arg Ile Thr Asp Val Leu Asp Gln Lys

Asn Tyr Val Glu Glu Leu Asn Arg His Leu Ser Cys Thr Val Gly Asp Leu Gln Thr Lys Ile Asp Gly Leu Glu Lys Thr Asn Ser Lys Leu Gln Glu Glu Leu Ser Ala Ala Thr Asp Arg Ile Cys Ser Leu Gln Glu Glu Gln Gln Gln Leu Arg Glu Gln Asn Glu Leu Ile Arg Glu Arg Ser Glu Lys Ser Val Glu Ile Thr Lys Gln Asp Thr Lys Val Glu Leu Glu Thr Tyr Lys Gln Thr Arg Gln Gly Leu Asp Glu Met Tyr Ser Asp Val Trp Lys Gln Leu Lys Glu Glu Lys Lys Val Arg Leu Glu Leu Glu Lys Glu Leu Glu Leu Gln Ile Gly Met Lys Thr Glu Met Glu Ile Ala Met Lys Leu Leu Glu Lys Asp Thr His Glu Lys Gln Asp Thr Leu Val Ala Leu Arg Gln Gln Leu Glu Glu Val Lys Ala Ile Asn Leu Gln Met Phe His Lys Ala Gln Asn Ala Glu Ser Ser Leu Gln Gln Lys Asn Glu Ala Ile Thr Ser Phe Glu Gly Lys Thr Asn Gln Val Met Ser Ser Met Lys Gln Met Glu Glu Arg Leu Gln His Ser Glu Arg Ala Arg Gln Gly Ala Glu Glu Arg Ser His Lys Leu Gln Gln Glu Leu Gly Gly Arg Ile Gly Ala Leu Gln Leu Gln Leu Ser Gln Leu His Glu Gln Cys Ser Ser Leu Glu Lys Glu Leu Lys Ser Glu Lys Glu Gln Arg Gln Ala Leu Gln Arg Glu 55 Leu Gln His Glu Lys Asp Thr Ser Ser Leu Leu Arg Met Glu Leu Gln Gln Val Glu Gly Leu Lys Lys Glu Leu Arg Glu Leu Gln Asp Glu Lys Ala Glu Leu Gln Lys Ile Cys Glu Glu Glu Glu Gln Ala Leu Gln Glu
595 600 605 Met Gly Leu His Leu Ser Gln Ser Lys Leu Lys Met Glu Asp Ile Lys Glu Val Asn Gln Ala Leu Lys Gly His Ala Trp Leu Lys Asp Asp Glu . . Ala Thr His Cys Arg Gln Cys Glu Lys Glu Phe Ser Ile Ser Arg Arg Lys His His Cys Arg Asn Cys Gly His Ile Phe Cys Asn Thr Cys Ser Ser Asn Glu Leu Ala Leu Pro Ser Tyr Pro Lys Pro Val Arg Val Cys Asp Ser Cys His Thr Leu Leu Gln Arg Cys Ser Ser Thr Ala Ser

<210> 192
<211> 331
<212> PRT

<213> Homo sapien

<400> 192 Arg Ala Gly Ala Ser Ala Met Ala Leu Arg Lys Glu Leu Leu Lys Ser - 1 10 Ile Trp Tyr Ala Phe Thr Ala Leu Asp Val Glu Lys Ser Gly Lys Val Ser Lys Ser Gln Leu Lys Val Leu Ser His Asn Leu Tyr Thr Val Leu 40 His Ile Pro His Asp Pro Val Ala Leu Glu Glu His Phe Arg Asp Asp 55 60 Asp Asp Gly Pro Val Ser Ser Gln Gly Tyr Met Pro Tyr Leu Asn Lys 70 80 Tyr Ile Leu Asp Lys Val Glu Glu Gly Ala Phe Val Lys Glu His Phe 85 90* Asp Glu Leu Cys Trp Thr Leu Thr Ala Lys Lys Asn Tyr Arg Ala Asp 100 105 Ser Asn Gly Asn Ser Met Leu Ser Asn Gln Asp Ala Phe Arg Leu Trp - 115 . 120 125 Cys Leu Phe Asn Phe Leu Ser Glu Asp Lys Tyr Pro Leu Ile Met Val 135 140 Pro Asp Glu Val Glu Tyr Leu Leu Lys Lys Val Leu Ser Ser Met Ser 145 150 155 . 160 Leu Glu Val Ser Leu Gly Glu Leu Glu Glu Leu Leu Ala Gln Glu Ala 165 170 175 🐇 🧻 Gln Val Ala Gln Thr Thr Gly Gly Leu Ser Val Trp Gln Phe Leu Glu 180 190 185 Leu Phe Asn Ser Gly Arg Cys Leu Arg Gly Val Gly Arg Asp Thr Leu 195 200 205 Ser Met Ala Ile His Glu Val Tyr Gln Glu Leu Ile Gln Asp Val Leu 210 215 220 Lys Gln Gly Tyr Leu Trp Lys Arg Gly His Leu Arg Arg Asn Trp Ala 225 230 235 Glu Arg Trp Phe Gln Leu Gln Pro Ser Cys Leu Cys Tyr Phe Gly Ser 245 250 255 Glu Glu Cys Lys Glu Lys Arg Gly Ile Ile Pro Leu Asp Ala His Cys 260 265 270 Cys Val Glu Val Leu Pro Asp Arg Asp Gly Lys Arg Cys Met Phe Cys 275 280 Val Lys Thr Ala Thr Arg Thr Tyr Glu Met Ser Ala Ser Asp Thr Arg .295 300 Gln Arg Gln Glu Trp Thr Ala Ala Ile Gln Met Ala Ile Arg Leu Gln 310 315 Ala Glu Gly Lys Thr Ser Leu His Lys Asp Leu 325

<210> 193 <211> 475 <212> PRT <213> Homo sapien

<400> 193

Lys Asn Ser Pro Leu Leu Ser Val Ser Ser Gln Thr Ile Thr Lys Glu

1 5 10 15

Asn Asn Arg Asn Val His Leu Glu His Ser Glu Gln Asn Pro Gly Ser

			20					25					30		
Ser	Ala	Gly 35	Asp	Thr	Ser	Ala	Ala 40	His	Gln	Val	Val	Leu 45	Gly	Glu	Asn
Leu	Ile 50	Ala	Thr	Ala	Leu	Cys 55	Leu	Ser	Gly	Ser	Gly 60	Ser	Gln	Ser	Asp
Leu 65	Lys	Asp	Val	Ala	Ser 70	Thr	Ala	Gly	Glü	Glu 75	Gly	Asp	Thr	Ser	Leu 80
	Glu	Ser	Leu	His 85	Pro	Val	Thr	Arg	Ser 90	Leu	Lys	Ala	Gly	Cys 95	His
Thr	Lys	Gln	Leu 100		Ser	Arg	Asn	Cys 105	Ser	Glu	Glu	Lys	Ser	Pro	Gln
Thr	Ser	Ile 115		Lys	Glu	Gly	Asn 120	Arg	Asp	Thr	Ser	Leu 125	Asp	Phe	Arg
Pro	Val		Ser	Pro	Ala	Asn 135		Va1	Glu	Gly	Val		Val	Asp	Gln
Asp 145	Asp	Asp	Gln	Asp	Ser 150		Ser	Leu	Lys	Leu 155		Gln	Asn	Ile	Ala 160
	Gln	Thr	qeA	Phe 165		Thr	Ala	Asp	Ser 170		Val	Àsn	Thr	Asp 175	
Asp	Ile	Glu	Lys 180		Leu	Asp	Lys	Met 185	Met	Thr	Glu	Arg	Thr 190		Leu
Lys	Glu	Arg 195		Gln	Glu	Val	Leu 200	Asp	Lys	Gln	Arg	Gln 205	Val	Glu	Asn
Gln	Leu 210	Gln	Val	Gln		Lys 215	Gln	Leu	Gln		Arg 220	Arg	Glu	Glu	Glu
Met 225	Lys	Asn	His	Gln	Glu 230	Ile	Leu	Lys	Ala	Ile 235	Gln	Asp	Val	Thr	11e 240
Lys	Arg	Glu	Glu	Thr 245	Lys	Lys	Lys	Ile	Glu 250	Lys	Glu	Lys	Lys	Glu 255	Phe
Leu	Gln	Lys	Glu 260	Gln	Asp	Leu	Lys	Ala 265	Glu	Ile	Glu	Lys	Leu 270	Cys	Glu a:
		275					280	·				285			
Gln	Asp 290	Gly	Glu	Ilė		Arg 295	Asn	Ile	Met	Glu	Glu 300	Thr	Glu	Arg	Ala
Trp 305	Lys	Ala	Glu	Ile	Leu 310	Ser	Leu	Glu	Ser	Arg 315	Lys	Glu	Leu	Leu	Val 320
Leu	Lys	Leu	Glu	Glu 325	Ala	Glu	Lys	Glu	Ala 330	Glu	Leu	His	Leu	Thr 335	Tyr
Leu	Lys	Ser	Thr 340	Pro	Pro	Thr	Leu	Glu 345		Val	Arg	Ser	Lys 350	Gln	Glu
Trp	Glu	Thr 355	Arg	Leu	Asn	Gly	Val 360	Arg	Ile	Met	Lys	Lys 365	Asn	Val	Arg
.Asp	Gln 370	Phe	Asn	Ser	His	Ile 375	Gln	Leu	Val	Arg	Asn 380	Gly	Ala	Lys	Leu
Ser 385	Ser	Leu	Pro		Ile 390	Pro	Thr	Pro	Thr	Leu 395	Pro	Pro	Pro	Pro	Ser 400
Glu	Thr	Asp		Met 405	Leu	Gln	Val	Phe	Gln 410	Pro	Ser	Pro	Ser	Leu 415	Ala
Pro	Arg	Met	Pro 420	Phe	Ser	Ile	Gly	Gln 425	Val	Thr	Met	Pro	Met 430	Val	Met
Pro	Ser	Ala 435	Asp	Pro	Arg	Ser	Leu 440	Ser	Phe	Pro	Ile	Leu 445	Asn	Pro	Ala
Leu	Ser 450	Gln	Pro	Ser	Gln	Pro 455	Ser	Ser	Pro	Leu	Pro 460	Gly	Ser	His	Gly

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Arg Asn Ser Pro Gly Leu Gly Ser Leu Val Ser
465 470 475
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<210> 194 <211> 241 <212> PRT <213> Homo sapien

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<210> 195 <211> 138 <212> PRT <213> Homo sapien

<400> 195

50 55 60 Gln Gln Glu His Ile His Glu Leu Gln Glu Leu Lys Asp Gln Leu Glu 70 Gln Gln Leu Gln Gly Leu His Arg Lys Val Gly Glu Thr Ser Leu Leu 85 90 Leu Ser Gln Arg Glu Gln Glu Ile Val Val Leu Gln Gln Gln Leu Gln 100 105 Glu Ala Arg Glu Gln Gly Glu Leu Lys Glu Gln Ser Leu Gln Ser Gln 115 120 Leu Asp Glu Ala Gln Arg Ala Leu Ala Gln 130 135 <210> 196 <211> 102

<400> 196

<212> PRT < <213> Homo sapien

Met Ser Lys Arg Lys Ala Pro Gln Glu Thr Leu Asn Gly Gly Ile Thr 5 10 15 Asp Met Leu Thr Glu Leu Ala Asn Phe Glu Lys Asn Val Ser Gln Ala A 201 25 30 Ile His Lys Tyr Asn Ala Tyr Arg Lys Ala Ala Ser Val Ile Ala Lys 35 40 45... Tyr Pro His Lys Ile Lys Ser Gly Ala Glu Ala Lys Lys Leu Pro Gly 50 55 60 Val Gly Thr Lys Ile Ala Glu Lys Ile Asp Glu Phe Leu Ala Thr Gly 65 70 75 Lys Leu Arg Lys Leu Glu Lys Ile Arg Gln Asp Asp Thr Ser Ser Ser 90 Ile Asn Phe Leu Thr Arg 100

<210> 197 <211> 138 <212> PRT <213> Homo sapien

<400> 197

Glu Ala Asn Glu Val Thr Asp Ser Ala Tyr Met Gly Ser Glu Ser Thr 10 15 Tyr Ser Glu Cys Glu Thr Phe Thr Asp Glu Asp Thr Ser Thr Leu Val 30 25 His Pro Glu Leu Gln Pro Glu Gly Asp Ala Asp Ser Ala Gly Gly Ser 35 40 45 Ala Val Pro Ser Glu Cys Leu Asp Ala Met Glu Glu Pro Asp His Gly 55 60 Ala Leu Leu Leu Pro Gly Arg Pro His Pro His Gly Gln Ser Val ··· 70 75 Ile Thr Val Ile Gly Gly Glu Glu His Phe Glu Asp Tyr Gly Glu Gly 85 90 Ser Glu Ala Glu Leu Ser Pro Glu Thr Leu Cys Asn Gly Gln Leu Gly 100 110 Cys Ser Asp Pro Ala Phe Leu Thr Pro Ser Pro Thr Lys Arg Leu Ser 115 120

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Ser Lys Lys Val Ala Arg Tyr Leu His Gln
   130
                       135
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<210> 198

<211> 100

<212> PRT

<213> Homo sapien

<400> 198

Met Gly Asp Val Lys Asn Phe Leu Tyr Ala Trp Cys Gly Lys Arg Lys 10 Met Thr Pro Ser Tyr Glu Ile Arg Ala Val Gly Asn Lys Asn Arg Gln 25 Lys Phe Met Cys Glu Val Gln Val Glu Gly Tyr Asn Tyr Thr Gly Met **35** .. 40 Gly Asn Ser Thr Asn Lys Lys Asp Ala Gln Ser Asn Ala Ala Arg Asp 55 ...60 Phe Val Asn Tyr Leu Val Arg Ile Asn Glu Ile Lys Ser Glu Glu Val 80 ~ 70 75 Pro Ala Phe Gly Val Ala Ser Pro Pro Pro Leu Thr Asp Thr Pro Asp 85 90

Thr Thr Ala Asn 100

<210> 199

∘ <212> PRT :

<213> Homo sapien

<400> 199

Met Val Lys Glu Thr Thr Tyr Tyr Asp Val Leu Gly Val Lys Pro Asn 10 Ala Thr Gln Glu Glu Leu Lys Lys Ala Tyr Arg Lys Leu Ala Leu Lys 25 20 Tyr His Pro Asp Lys Asn Pro Asn Glu Gly Glu Lys Phe Lys Gln Ile 40 Ser Gln Ala Tyr Glu Val Leu Ser Asp Ala Lys Lys Arg Glu Leu Tyr 55 Asp Lys Gly Gly Glu Gln Ala Ile Lys Glu Gly Gly Ala Gly Gly Gly 70 75 Phe Gly Ser Pro Met Asp Ile Phe Asp Met Phe Phe Gly Gly Gly 85 90 Arg Met Gln Arg Glu Arg Arg Gly Lys Asn Val Val His Gln Leu Ser 100 110 105 Val Thr Leu Glu Asp Leu Tyr Asn Gly Ala Thr Arg Lys Leu Ala 120 125

<210> 200

<211> 90

<212> PRT

<213> Homo sapien

<400> 200

Met Ala Cys Pro Leu Asp Gln Ala Ile Gly Leu Leu Val Ala Ile Phe 10

<210> 201 <211> 120 <212> PRT <213> Homo sapien

<400> 201 Met Glu Thr Pro Ser Gln Arg Arg Ala Thr Arg Ser Gly Ala Gln Ala Ser Ser Thr Pro Leu Ser Pro Thr Arg Ile Thr Arg Leu Gln Glu Lys Glu Asp Leu Gln Glu Leu Asn Asp Arg Leu Ala Val Tyr Ile Asp Arg Val Arg Ser Leu Glu Thr Glu Asn Ala Gly Leu Arg Leu Arg Ile Thr 55 Glu Ser Glu Glu Val Val Ser Arg Glu Val Ser Gly Ile Lys Ala Ala 70 75 Tyr Glu Ala Glu Leu Gly Asp Ala Arg Lys Thr Leu Asp Ser Val Ala 90 85 Lys Glu Arg Ala Arg Leu Gln Leu Glu Leu Ser Lys Val Arg Glu Glu 100 105 Phe Lys Glu Leu Lys Ala Arg Asn 115

<210> 202 <211> 177 <212> PRT <213> Homo sapien

<210> 203 <211> 164 <212> PRT <213> Homo sapien

<400> 203

Met Arg Leu Ala Val Gly Ala Leu Leu Val Cys Ala Val Leu Gly Leu 5 -- 10 15 Cys Leu Ala Val Pro Asp Lys Thr Val Arg Trp Cys Ala Val Ser Glu 20 . 25 His Glu Ala Thr Lys Cys Gln Ser Phe Arg Asp His Met Lys Ser Val 40 IIe Pro Ser Asp Gly Pro Ser Val Ala Cys Val Lys Lys Ala Ser Tyr 60 Leu Asp Cys Ile Arg Ala Ile Ala Ala Asn Glu Ala Asp Ala Val Thr 70 75 Leu Asp Ala Gly Leu Val Tyr Asp Ala Tyr Leu Ala Pro Asn Asn Leu 90 Lys Pro Val Val Ala Glu Phe Tyr Gly Ser Lys Glu Asp Pro Gln Thr 100 105 Phe Tyr Tyr Ala Val Ala Val Val Lys Lys Asp Ser Gly Phe Gln Met 115 120 Asn Gln Leu Arg Gly Lys Lys Ser Cys His Thr Gly Leu Gly Arg Ser 135 140 Ala Gly Trp Asn Ile Pro Ile Gly Leu Leu Tyr Cys Asp Leu Pro Glu 155 Pro Arg Lys Pro

<210> 204 <211> 241 <212> PRT <213> Homo sapien

<400> 204

 Met
 Ser
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 Glu
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 Gly
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 Ala
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 Pro
 Glu
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 Ser
 Ile
 Arg
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 Tyr
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Gln Leu Ile Tyr Glu Ser Ala Ile Thr Cys Glu Tyr Leu Asp Glu Ala 85 90 Tyr Pro Gly Lys Lys Leu Leu Pro Asp Asp Pro Tyr Glu Lys Ala Cys 100 105 Gln Lys Met Ile Leu Glu Leu Phe Ser Lys Val Pro Ser Leu Val Gly 115 125 120 Ser Phe Ile Arg Ser Gln Asn Lys Glu Asp Tyr Asp Gly Leu Lys Glu 130 135 140 Glu Phe Arg Lys Glu Phe Thr Lys Leu Glu Glu Val Leu Thr Asn Lys 150 155 Lys Thr Thr Phe Phe Gly Gly Asn Ser Ile Ser Met Ile Asp Tyr Leu 165 170 Ile Trp Pro Trp Phe Glu Arg Leu Glu Ala Met Lys Leu Asn Glu Cys 185 Val Asp His Thr Pro Lys Leu Lys Leu Trp Met Ala Ala Met Lys Glu 200 Asp Pro Thr Val Ser Ala Leu Leu Thr Ser Glu Lys Asp Trp Gln Gly 220 Phe Leu Glu Leu Tyr Leu Gln Asn Ser Pro Glu Ala Cys Asp Tyr Gly 225 230 235 Leu

<210> 205 <211> 160 <212> PRT <213> Homo sapien

<400> 205

Met Gln Ile Phe Val Lys Thr Leu Thr Gly Lys Thr Ile Thr Leu Glu 5 10 Val Glu Pro Ser Asp Thr Ile Glu Asn Val Lys Ala Lys Ile Gln Asp 25 Lys Glu Gly Ile Pro Pro Asp Gln Gln Arg Leu Ile Phe Ala Gly Lys 35 40 Gln Leu Glu Asp Gly Arg Thr Leu Ser Asp Tyr Asn Ile Gln Lys Glu . 50 55 60 Ser Thr Leu His Leu Val Leu Arg Leu Arg Gly Gly Met Gln Ile Phe 70 75 Val Lys Thr Leu Thr Gly Lys Thr Ile Thr Leu Glu Val Glu Pro Ser 85 90 Asp Thr Ile Glu Asn Val Lys Ala Lys Ile Gln Asp Lys Glu Gly Ile 100 105 Pro Pro Asp Gln Gln Arg Leu Ile Phe Ala Gly Lys Gln Leu Glu Asp 120 Gly Arg Thr Leu Ser Asp Tyr Asn Ile Gln Lys Glu Ser Thr Leu His 130 140 Leu Val Leu Arg Leu Arg Gly Gly Met Gln Ile Phe Val Lys Thr Leu 145 150 155 160

<210> 206

<211> 197

<212> PRT

<213> Homo sapien

Thr Ser Pro Ser Glu Ala Cys Ala Pro Leu Leu Ile Ser Leu Ser Thr 10 Leu Ile Tyr Asn Gly Ala Leu Pro Cys Gln Cys Asn Pro Gln Gly Ser 20 25 Leu Ser Ser Glu Cys Asn Pro His Gly Gly Gln Cys Leu Cys Lys Pro 40 Gly Val Val Gly Arg Arg Cys Asp Leu Cys Ala Pro Gly Tyr Tyr Gly 55 Phe Gly Pro Thr Gly Cys Gln Gly Ala Cys Leu Gly Cys Arg Asp His Thr Gly Gly Glu His Cys Glu Arg Cys Ile Ala Gly Phe His Gly Asp Pro Arg Leu Pro Tyr Gly Gly Gln Cys Arg Pro Cys Pro Cys Pro Glu 105 Gly Pro Gly Ser Gln Arg His Phe Ala Thr Ser Cys His Gln Asp Glu 120 115 125 Tyr Ser Gln Gln Ile Val Cys His Cys Arg Ala Gly Tyr Thr Gly Leu 135 Arg Cys Glu Ala Cys Ala Pro Gly His Phe Gly Asp Pro Ser Arg Pro 150 155 Gly Gly Arg Cys Gln Leu Cys Glu Cys Ser Gly Asn Ile Asp Pro Met 170 165 Asp Pro Asp Ala Cys Asp Pro His Thr Gly Gln Cys Leu Arg Cys Leu His His Thr Glu Gly 195

<210> 207 <211> 175 <212> PRT <213> Homo sapien

<400> 207

Ile Ile Arg Gln Gln Gly Leu Ala Ser Tyr Asp Tyr Val Arg Arg Arg 10 Leu Thr Ala Glu Asp Leu Phe Glu Ala Arg Ile Ile Ser Leu Glu Thr Tyr Asn Leu Leu Arg Glu Gly Thr Arg Ser Leu Arg Glu Ala Leu Glu 40 Ala Glu Ser Ala Trp Cys Tyr Leu Tyr Gly Thr Gly Ser Val Ala Gly 55 Val Tyr Leu Pro Gly Ser Arg Gln Thr Leu Ser Ile Tyr Gln Ala Leu 70 Lys Lys Gly Leu Leu Ser Ala Glu Val Ala Arg Leu Leu Glu Ala 90 Gln Ala Ala Thr Gly Phe Leu Leu Asp Pro Val Lys Gly Glu Arg Leu 105 Thr Val Asp Glu Ala Val Arg Lys Gly Leu Val Gly Pro Glu Leu His 120 Asp Arg Leu Leu Ser Ala Glu Arg Ala Val Thr Gly Tyr Arg Asp Pro 135 140 Tyr Thr Glu Gln Thr Ile Ser Leu Phe Gln Ala Met Lys Lys Glu Leu 150 155 Ile Pro Thr Glu Glu Ala Leu Arg Leu Trp Met Pro Ser Trp Pro

165 170 175

<210> 208 <211> 177 <212> PRT <213> Homo sapien

<400> 208

Met Ala Ala Gly Val Glu Ala Ala Ala Glu Val Ala Ala Thr Glu Ile 1 10 Lys Met Glu Glu Glu Ser Gly Ala Pro Gly Val Pro Ser Gly Asn Gly 25 Ala Pro Gly Pro Lys Gly Glu Gly Glu Arg Pro Ala Gln Asn Glu Lys ∂; **40** Arg Lys Glu Lys Asn Ile Lys Arg Gly Gly Asn Arg Phe Glu Pro Tyr 55 - 60 Ala Asn Pro Thr Lys Arg Tyr Arg Ala Phe Ile Thr Asn Ile Pro Phe 70 Asp Val Lys Trp Gln Ser Leu Lys Asp Leu Val Lys Glu Lys Val Gly 85 (will be 90. Glu Val Thr Tyr Val Glu Leu Leu Met Asp Ala Glu Gly Lys Ser Arg 100 ·· 105 -Gly Cys Ala Val Val Glu Phe Lys Met Glu Glu Ser Met Lys Lys Ala 120 115 125 Ala Glu Val Leu Asn Lys His Ser Leu Ser Gly Arg Pro Leu Lys Val 135 140 Lys Glu Asp Pro Asp Gly Glu His Ala Arg Arg Ala Met Gln Lys Val 150 155 Met Ala Thr Thr Gly Gly Met Gly Met Gly Pro Gly Gly Pro Gly Met 165 170 175 Ile

<210> 209 <211> 196 <212> PRT <213> Homo sapien

<400> 209

Asp Leu Gln Asp Met Phe Ile Val His Thr Ile Glu Glu Ile Glu Gly 10 Leu Ile Ser Ala His Asp Gln Phe Lys Ser Thr Leu Pro Asp Ala Asp 25 Arg Glu Arg Glu Ala Ile Leu Ala Ile His Lys Glu Ala Gln Arg Ile 40 Ala Glu Ser Asn His Ile Lys Leu Ser Gly Ser Asn Pro Tyr Thr Thr 55 60 te-Val Thr Pro Gln Ile Ile Asn Ser Lys Trp Glu Lys Val Gln Gln Leu 70 75 Val Pro Lys Arg Asp His Ala Leu Leu Glu Glu Gln Ser Lys Gln Gln 90 Ser Asn Glu His Leu Arg Arg Gln Phe Ala Ser Gln Ala Asn Val Val 105 Gly Pro Trp Ile Gln Thr Lys Met Glu Glu Ile Gly Arg Ile Ser Ile 115 125

1 138E 1 7

1989 1984 1983

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Glu Met Asn Gly Thr Leu Glu Asp Gln Leu Ser His Leu Lys Gln Tyr
                     135
Glu Arg Ser Ile Val Asp Tyr Lys Pro Asn Leu Asp Leu Leu Glu Gln
                                      155 ..
Gln His Gln Leu Ile Gln Glu Ala Leu Ile Phe Asp Asn Lys His Thr
              165
                                 170
Asn Tyr Thr Met Glu His Ile Arg Val Gly Trp Glu Gln Leu Leu Thr
          180
Thr Ile Ala Arg
        195
      <210> 210
      <211> 156
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Lys Leu Thr Ile Glu Ser Thr Pro Phe Asn Val Ala Glu Gly Lys Glu
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Val Leu Leu Leu Ala His Asn Leu Pro Gln Asn Arg Ile Gly Tyr Ser
  20 25
                                                 30
Trp Tyr Lys Gly Glu Arg Val Asp Gly Asn Ser Leu Ile Val Gly Tyr
  35
                         40
Val Ile Gly Thr Gln Gln Ala Thr Pro Gly Pro Ala Tyr Ser Gly Arg
50
60
Glu Thr Ile Tyr Pro Asn Ala Ser Leu Leu Ile Gln Asn Val Thr Gln
                   70.
Asn Asp Thr Gly Phe Tyr Thr Leu Gln Val Ile Lys Ser Asp Leu Val
               85
                                  90
Asn Glu Glu Ala Thr Gly Gln Phe His Val Tyr Pro Glu Leu Pro Lys
          100
                             105
Pro Ser Ile Ser Ser Asn Asn Ser Asn Pro Val Glu Asp Lys Asp Ala
     115
                         120
Val Ala Phe Thr Cys Glu Pro Glu Val Gln Asn Thr Thr Tyr Leu Trp
                                         140
                     135
Trp Val Asn Gly Gln Ser Leu Pro Val Ser Pro Lys
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85

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<400> 212

Glu Lys Gln Lys Asn Lys Glu Phe Ser Gln Thr Leu Glu Asn Glu Lys 10 Asn Thr Leu Leu Ser Gln Ile Ser Thr Lys Asp Gly Glu Leu Lys Met 25 30 Leu Gln Glu Glu Val Thr Lys Met Asn Leu Leu Asn Gln Gln Ile Gln 40 45 Glu Glu Leu Ser Arg Val Thr Lys Leu Lys Glu Thr Ala Glu Glu Glu 55 60 Lys Asp Asp Leu Glu Glu Arg Leu Met Asn Gln Leu Ala Glu Leu Asn 70 Gly Ser Ile Gly Asn Tyr Cys Gln Asp Val Thr Asp Ala Gln Ile Lys Asn Glu Leu Leu Glu Ser Glu Met Lys Asn Leu Lys Lys Cys Val Ser 100 105 Glu Leu Glu Glu Lys Gln Gln Leu Val Lys Glu Lys Thr Lys Val 120 Glu Ser Glu Ile Arg Lys Glu Tyr Leu Glu Lys Ile Gln Gly 140 135 ..

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<213> Homo sapien

<400>. 214 Val Met Arg Val Asp Phe Asn Val Pro Met Lys Asn Asn Gln Ile Thr 15 Page 15 Pag Asn Asn Gln Arg Ile Lys Ala Ala Val Pro Ser Ile Lys Phe Cys Leu 30.5 Asp Asn Gly Ala Lys Ser Val Val Leu Met Ser His Leu Gly Arg Pro (1),35 (1) (1) (1) (1) (2) (40 (1) (1) (1) (1) (1) (45 (1) (1) (2) (2) (1) Asp Gly Val Pro Met Pro Asp Lys Tyr Ser Leu Glu Pro Val Ala Val ο καταγοργ**55**η ο κητηρού και ο **60**μο ή το πολού Glu Leu Arg Ser Leu Leu Gly Lys Asp Val Leu Phe Leu Lys Asp Cys 5.70 pt. 18. 18. 18. 18. 175 to the street of a price 80 to Val Gly Pro Glu Val Glu Lys Ala Cys Ala Asn Pro Ala Ala Gly Ser Val Ile Leu Clu Asn Leu Arg Phe His Val Glu Glu Glu Gly Lys
100 105 110 Gly Lys Asp Ala Ser Gly Asn Lys Val Lys Ala Glu Pro Ala Lys Ile 115

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নার স্কৃতি । তার্মার্কিক কোনো কর্মান কার্ত্তার । তার্কালিক কোনার সুক্রাই জেন্তুর ইতিকার কার্ত্তার ব্যবহার বিভাগিক কোনার ভারতি ইতিকার হয়

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WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁶ : C12N 15/12, A61K 38/17, C07K 14/47, A3 16/18, A61K 35/14	(11) International Publication Number: WO 99/38973 (43) International Publication Date: 5 August 1999 (05.08.99)
09/015,022 28 January 1998 (28.01.98) 09/040,828 18 March 1998 (18.03.98) 09/040,831 18 March 1998 (18.03.98) 09/122,192 23 July 1998 (23.07.98) 09/122,191 23 July 1998 (23.07.98)	BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TI, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Burasian patent (AM, AZ, BY, KG, KZ, MD, RU, TI, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published With international search report.
 (72) Inventors: REED, Steven, G.; 2843 – 122nd Place N Bellevue, WA 98005 (US). LODES, Michael, J.; 922 36th Avenue S.W., Seattle, WA 98126 (US). FRUDAK Tony, N.; P.O. Box 99232, Seattle, WA 99232-0232 (UMOHAMATH, Raodoh; 4205 South Morgan, Seattle, V 98118 (US). (74) Agents: MAKI, David, J. et al.; Seed and Berry L 6300 Columbia Center, 701 Fifth Avenue, Seattle, V 98104-7092 (US). 	9 December 1999 (09.12.99) SS. VA

(57) Abstract

Compounds and methods for treating lung cancer are provided. The inventive compounds include polypeptides containing at least a portion of a lung tumor protein. Vaccines and pharmaceutical compositions for immunotherapy of lung cancer comprising such polypeptides, or polynucleotides encoding such polypeptides, are also provided, together with polynucleotides for preparing the inventive polypeptides.

INTERNATIONAL SEARCH REPORT

Inte Sonal Application No
PCT/US 99/01642

CLASSIF	C12N15/12 A61K38/17 C07K14/4	7 C07K16/18	A61K	35/14
ecording to	International Patent Classification (IPC) or to both national classification	ion and IPC		
	SEARCHED		· · · · · ·	
	cumentation searched (classification system followed by classification C12N C12O A61K C07K	1 вупізов)		
	ion searched other than minimum documentation to the extent that su	oh documents are included	in the fields see	rched
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DOCUME	ENTS CONSIDERED TO BE RELEVANT			
ategory *	Citation of document, with indication, where appropriate, of the rele	vant passages		Relevant to claim No.
	WO 96 30389 A (MILLENIUM PHARMACE	UTICALS,		1-60
	INC.; SHYJAN A.) 3 October 1996 see page 112 - page 127			
		.41	. 1	
١	WO 96 02552 A (CYTOCLONYL PHARMAC			1-60
	INC.; TORCZYNSKI R. ET AL.) 1 Feb 1996	ruary `	*	and the second
	see the whole document			· .
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`	YOU L ET AL.: "Identification of growth response gene-1 (Egr-1) as			1,2,4-/
	phorbol myristate-induced gene in	ı lung ·		
	cancer cells by differential mRN/	\ display"		
	AM. J. RESPIR. CELL MOL. BIOL., vol. 17, no. 5, November 1997,			
	pages 617-624, XP002106654			
	see page 618, left-hand column, p	paragraph		
	3			
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X Furt	her documents are listed in the continuation of box C.	X Patent family me	embers are listed	in sernex.
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2	21 June 1999	2	2 10 1999	
Name and	mailing address of the ISA	Authorized officer		
	European Patent Office, P.B. 5818 Patentinan 2 NL - 2280 HV Rijserijk	1		

INTERNATIONAL SEARCH REPORT

PCT/US 99/01642

Box	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)	
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:	•
. IV		
. (A)	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:	
	Remark: Although claims 16, 17, 24-26, 32, 33, 48-53 and 56-58 are	•
	directed to a method of treatment of the human/animal body	
	the search has been carried out and based on the alleged effects of the composition.	
. \Box		
س ۽	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such	
	an extent that no meaningful International Search can be carried out, specifically:	
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ı 🗌	Claims Nos.: because they are dependent claims and are not draited in accordance with the second and third sentences of Rule 6.4(a).	
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lox II	Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)	
This Int	ternational Searching Authority found multiple inventions in this international application, as follows:	
	ee FURTHER INFORMATION sheet	
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'· [As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.	
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² L_	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.	
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s. [As only some of the required additional search fees were timely paid by the applicant, this International Search Report	
L	covers only those claims for which fees were paid, specifically claims Nos.:	
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a X	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:	
-	Tobalish with a resident and the manufacture of the country and covered by commentation.	
	see FURTHER INFORMATION sheet, subject 1.	•
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Roma	rk on Protest The additional search fees were accompanied by the applicant's protest	•
	No protest accompanied the payment of additional search fees.	

INTERNATIONAL SEARCH REPORT

Information on patent family members

Inte 'onal Application No PCT/US 99/01642

Patent document cited in search report		Publication date		atent family nember(s)	Publication date
WO 9630389	A	03-10-1996	US AU AU CA EP US	5633161 A 708746 B 5437896 A 2216717 A 0817792 A 5674739 A	27-05-1997 12-08-1999 16-10-1996 03-10-1996 14-01-1998 07-10-1997
WO 9602552	A	01 -0 2-1996	US AU AU BR CA EP JP US	5589579 A 700915 B 3359295 A 9508417 A 2195403 A 0804451 A 10503087 T 5773579 A	31-12-1996 14-01-1999 16-02-1996 18-11-1997 01-02-1996 05-11-1997 24-03-1998